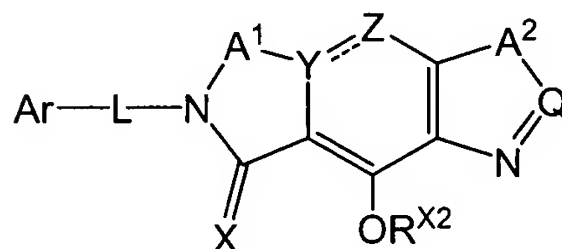


Amendments to the Claims

We claim:

- (Previously presented): A compound having the structure:

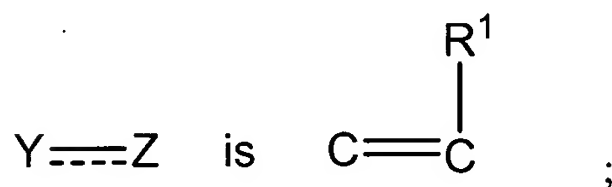


wherein:

A¹ is independently selected from C(R²)₂, CR²OR, CR²OC(=O)R, C(=O), C(=S), CR²SR, and C(=NR),

A² is independently selected from C(R²)₂-C(R³)₂, C(R²)=C(R³) and C(=O)C(R³)₂;

Q is CR⁴;



L is selected from a bond, O, S, S-S, S(=O), S(=O)₂, S(=O)₂NR, NR, N-OR, C₁-C₁₂ alkylene, C₁-C₁₂ substituted alkylene, C₂-C₁₂ alkenylene, C₂-C₁₂ substituted alkenylene, C₂-C₁₂ alkynylene, C₂-C₁₂ substituted alkynylene, C(=O)NH, OC(=O)NH, NHC(=O)NH, C(=O), C(=O)NH(CH₂)_n, or (CH₂CH₂O)_n, where n is optionally 1, 2, 3, 4, 5, or 6;

X is selected from O, S, NH, NR, N-OR, N-NR₂, N-CR₂OR and N-CR₂NR₂;

Ar is selected from (a) a C₃-C₁₂ carbocycle, C₃-C₁₂ substituted carbocycle, C₆-C₂₀ aryl, C₆-C₂₀ substituted aryl, C₂-C₂₀ heteroaryl, and C₂-C₂₀ substituted heteroaryl;

or (b) a saturated, unsaturated or aromatic ring or ring system having a mono- or bicyclic carbocycle or heterocycle containing 3 to 12 ring atoms;

R², R³ and R⁴ are each independently selected from H, F, Cl, Br, I, OH, -NH₂, -NH₃⁺, -NHR, -NR₂, -NR₃⁺, C₁-C₈ alkylhalide, carboxylate, sulfate, sulfamate, sulfonate, 5-7 membered ring sultam, C₁-C₈ alkylsulfonate, C₁-C₈ alkylamino, 4-dialkylaminopyridinium, C₁-C₈ alkylhydroxyl, C₁-C₈ alkylthiol, -SO₂R, -SO₂Ar, -SOAr, -SAr, -SO₂NR₂, -SOR, -CO₂R,

$-C(=O)NR_2$, 5-7 membered ring lactam, 5-7 membered ring lactone, $-CN$, $-N_3$, $-NO_2$, C_1-C_8 alkoxy, C_1-C_8 trifluoroalkyl, C_1-C_8 alkyl, C_1-C_8 substituted alkyl, C_3-C_{12} carbocycle, C_3-C_{12} substituted carbocycle, C_6-C_{20} aryl, C_6-C_{20} substituted aryl, C_2-C_{20} heteroaryl, and C_2-C_{20} substituted heteroaryl, polyethyleneoxy, phosphonate, phosphate, and a prodrug moiety;

when taken together on a single carbon, two R^2 or two R^3 may form a spiro ring;

R^1 is independently selected from CR_3 , $NRSO_2R$, $OC(=O)NR_2$, $OC(=O)R$, SR , H , F , Cl , Br , I , OH , $-NH_2$, $-NH_3^+$, $-NHR$, $-NR_2$, $-NR_3^+$, C_1-C_8 alkylhalide, carboxylate, sulfate, sulfamate, sulfonate, 5-7 membered ring sultam, C_1-C_8 alkylsulfonate, C_1-C_8 alkylamino, 4-dialkylaminopyridinium, C_1-C_8 alkylhydroxyl, C_1-C_8 alkylthiol, $-SO_2R$, $-SO_2Ar$, $-SOAr$, $-SAr$, $-SO_2NR_2$, $-SOR$, $-CO_2R$, $-C(=O)NR_2$, 5-7 membered ring lactam, 5-7 membered ring lactone, $-CN$, $-N_3$, $-NO_2$, C_1-C_8 alkoxy, C_1-C_8 trifluoroalkyl, C_1-C_8 alkyl, C_1-C_8 substituted alkyl, C_3-C_{12} carbocycle, C_3-C_{12} substituted carbocycle, C_6-C_{20} aryl, C_6-C_{20} substituted aryl, C_2-C_{20} heteroaryl, and C_2-C_{20} substituted heteroaryl, polyethyleneoxy, phosphonate, phosphate, and a prodrug moiety;

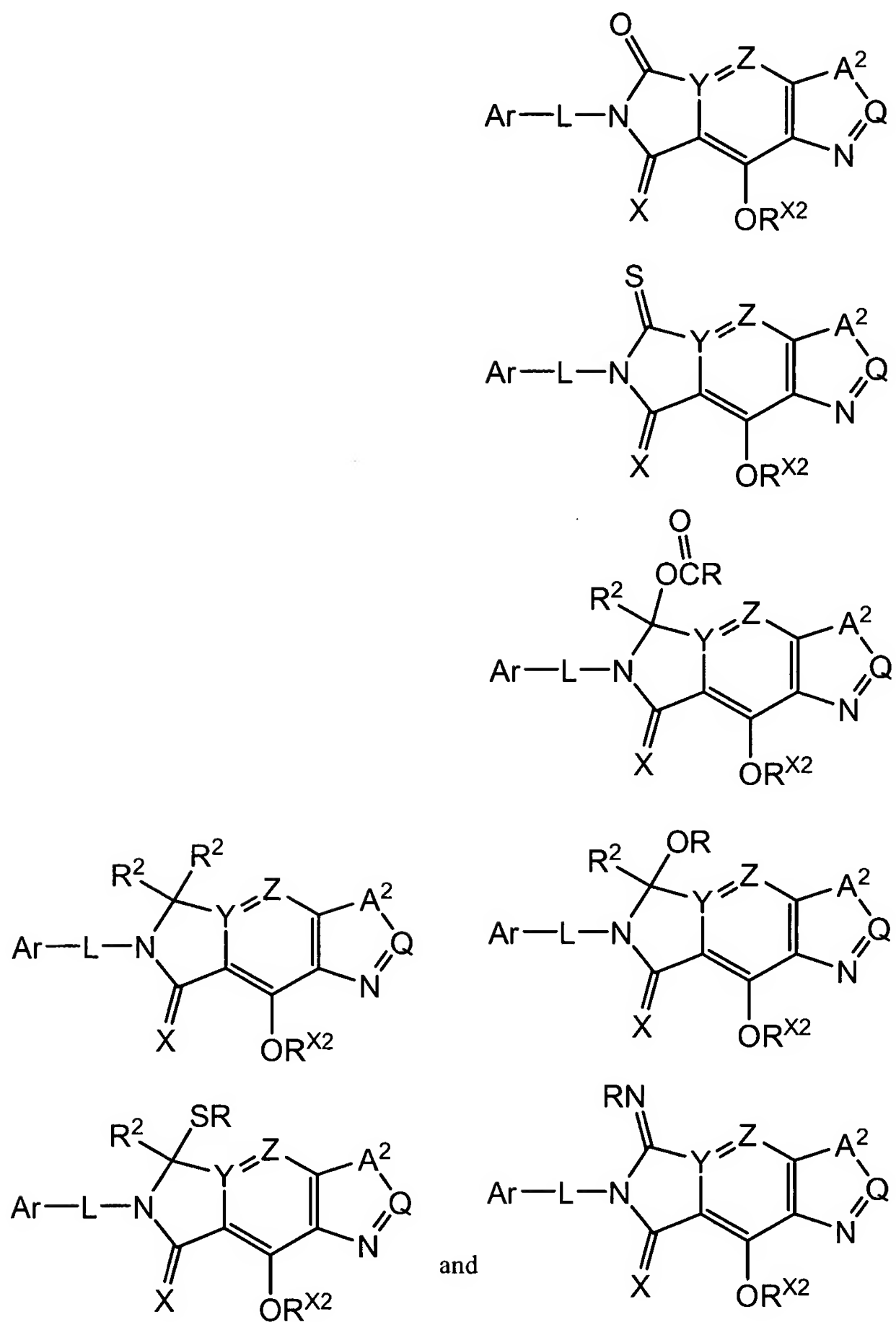
R is independently selected from H , C_1-C_8 alkyl, C_1-C_8 substituted alkyl, C_6-C_{20} aryl, C_6-C_{20} substituted aryl, C_2-C_{20} heteroaryl, and C_2-C_{20} substituted heteroaryl, polyethyleneoxy, phosphonate, phosphate, and a prodrug moiety;

R^{X2} is independently selected from H , C_1-C_8 alkyl, C_1-C_8 substituted alkyl, C_6-C_{20} aryl, C_6-C_{20} substituted aryl, C_2-C_{20} heteroaryl, and C_2-C_{20} substituted heteroaryl, polyethyleneoxy, phosphonate, phosphate, a prodrug moiety, and a protecting group;

and the tautomers, salts, solvates, resolved enantiomers and purified diastereomers thereof;

with the proviso that when $Y=Z$ is $C=C(OH)$, X is O , A^1 is $C(=O)$, A^2 is $C(R^2)=C(R^3)$, and Q is CH , then L is not a bond.

2. (Previously presented): A compound of claim 1 selected from the structures:

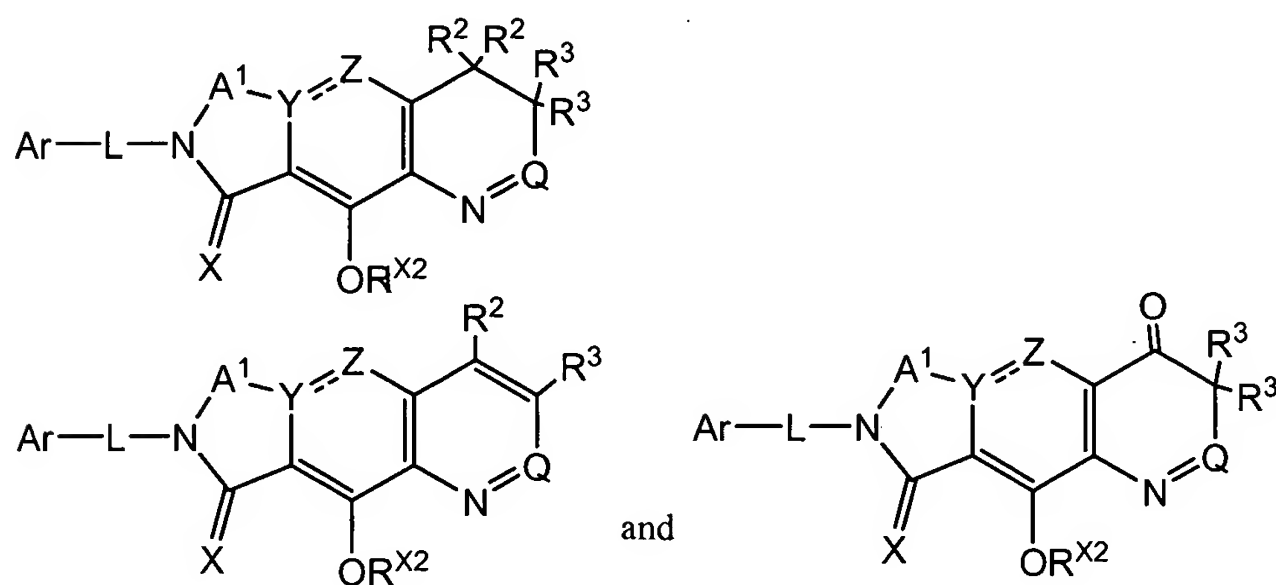


3. (Previously presented): A compound of claim 1 wherein A^1 is CH_2 .

4. (Previously presented): A compound of claim 1 wherein Ar is a saturated, unsaturated or aromatic ring or ring system having a mono- or bicyclic carbocycle or heterocycle containing 3 to 12 ring atoms.

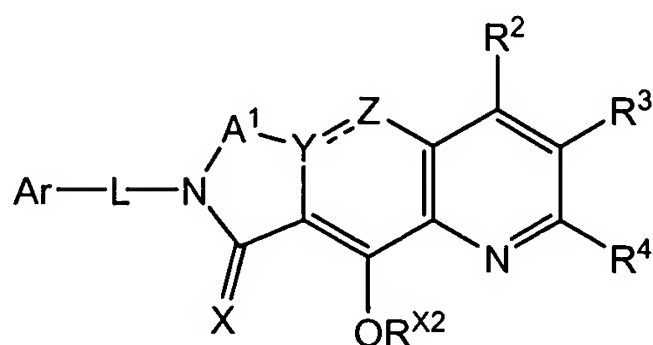
5. (Previously presented): A compound of claim 1 wherein R^1 is selected from R, OR, NR_2 , NHR, $NHSO_2R$ and $NRSO_2R$.

6. (Previously presented): A compound of claim 1 selected from the structures:

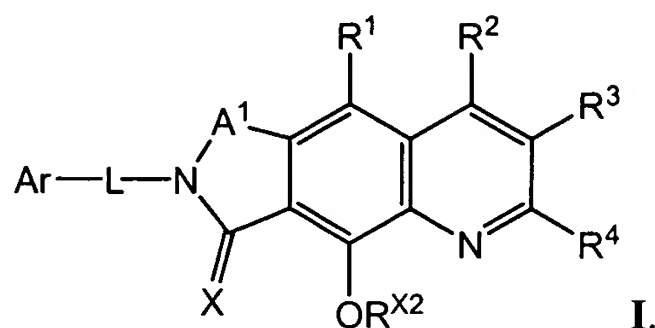


7. (Previously presented): A compound of claim 1 wherein R^{X2} is a protecting group selected from the group consisting of benzyhydryl ($CHPh_2$), trialkylsilyl (R_3Si), 2-trimethylsiloxyethyl, alkoxymethyl (CH_2OR), and ester ($C(=O)R$).

8. (Previously presented): A compound of claim 1 having the structure:

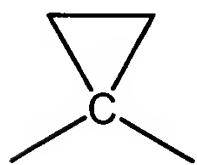


9. (Previously presented): A compound of claim 1 having Formula I:



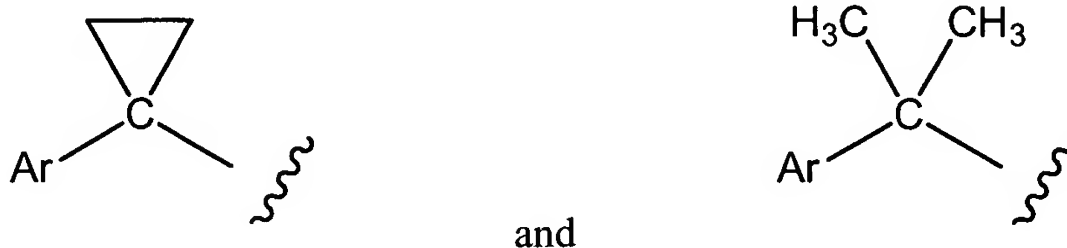
wherein R^{x2} is H and X is O.

10. (Previously presented): A compound of claim 1 wherein L is not a bond.
11. (Previously presented): A compound of claim 1 wherein R^4 is H.
12. (Previously presented): A compound of claim 1 wherein Ar is $C_6 - C_{20}$ substituted aryl.
13. (Previously presented): A compound of claim 1 having at least one phosphonate group.
14. (Previously presented): A compound of claim 1 wherein substituted alkyl, substituted alkylene, substituted alkyenylene, substituted alkynylene, substituted carbocycle, substituted aryl, and substituted heteroaryl are independently substituted with one or more substituents selected from F, Cl, Br, I, OH, $-NH_2$, $-NH_3^+$, $-NHR$, $-NR_2$, $-NR_3^+$, $C_1 - C_8$ alkylhalide, carboxylate, sulfate, sulfamate, sulfonate, 5-7 membered ring sultam, $C_1 - C_8$ alkylsulfonate, $C_1 - C_8$ alkylamino, 4-dialkylaminopyridinium, $C_1 - C_8$ alkylhydroxyl, $C_1 - C_8$ alkylthiol, $-SO_2R$, $-SO_2Ar$, $-SOAr$, $-SAr$, $-SO_2NR_2$, $-SOR$, $-CO_2R$, $-C(=O)NR_2$, 5-7 membered ring lactam, 5-7 membered ring lactone, $-CN$, $-N_3$, $-NO_2$, $C_1 - C_8$ alkoxy, $C_1 - C_8$ trifluoroalkyl, $C_1 - C_8$ alkyl, $C_3 - C_{12}$ carbocycle, $C_6 - C_{20}$ aryl, $C_2 - C_{20}$ heteroaryl, polyethyleneoxy, phosphonate, phosphate, and a prodrug moiety.
15. (Previously presented): The compound of claim 1 wherein A^1 is CH_2 , $C(CH_3)_2$,

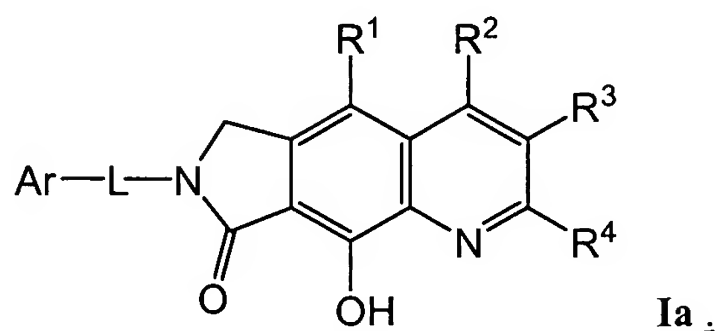


, $CH_2 - CH_2$, $C(CH_3)_2 - CH_2$, or $CH_2 - CH_2 - CH_2$.

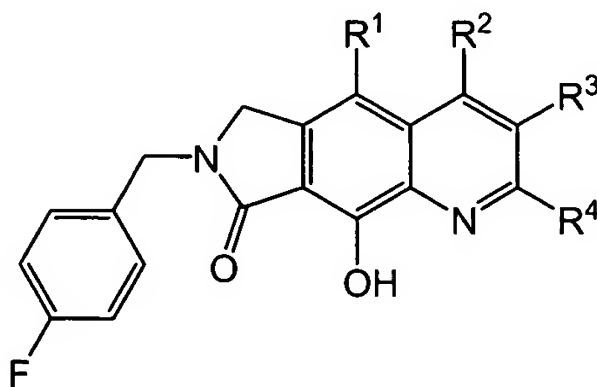
16. (Previously presented): The compound of claim 9 wherein X is O; L is CH₂; and Ar is substituted phenyl.
17. (Previously presented): The compound of claim 16 wherein Ar is 4-fluorophenyl.
18. (Previously presented): The compound of claim 9 wherein X is O; and R², R³ and R⁴ are each H.
19. (Previously presented): The compound of claim 9 wherein X is O; A¹ is CH₂; and R², R³ and R⁴ are each H.
20. (Previously presented): The compound of claim 1 wherein Ar-L is selected from the structures:



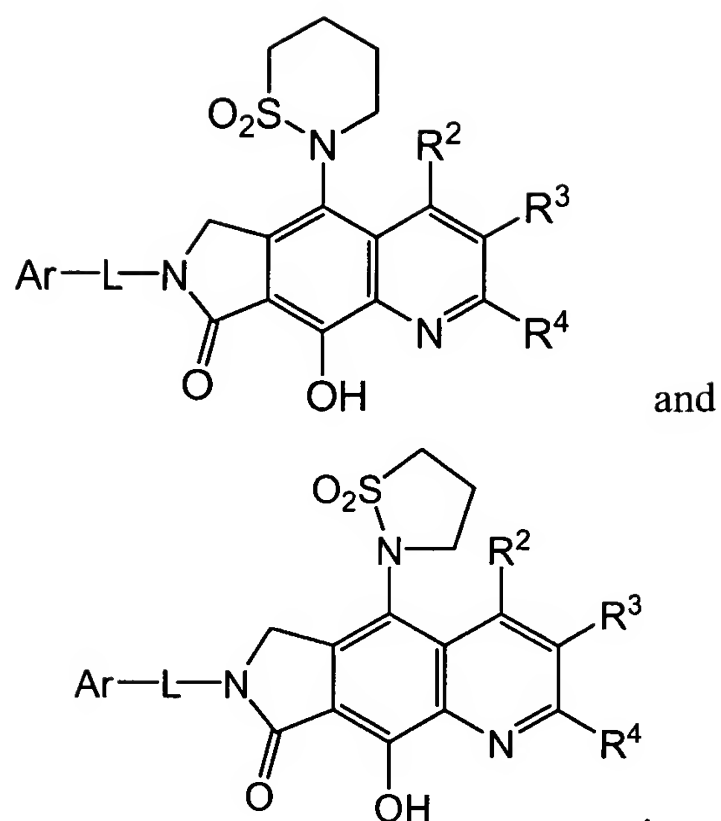
21. (Currently amended): A compound of claim 9 having Formula Ia:



22. (Previously presented): A compound of claim 9 having the structure:

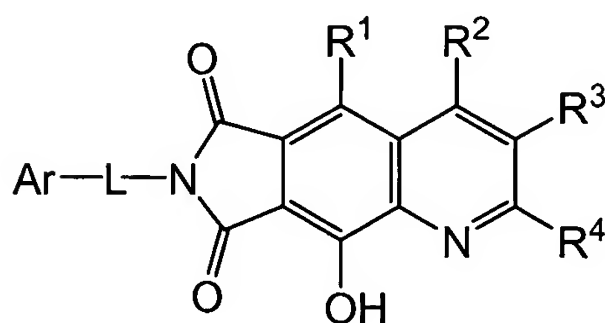


23. (Previously presented): A compound of claim 22 selected from the structures:



24. (Currently amended): A compound of claim 9 wherein ~~Ar-L~~ Ar-L is para-fluorobenzyl.

25. (Previously presented): A compound of claim 9 having the structure:



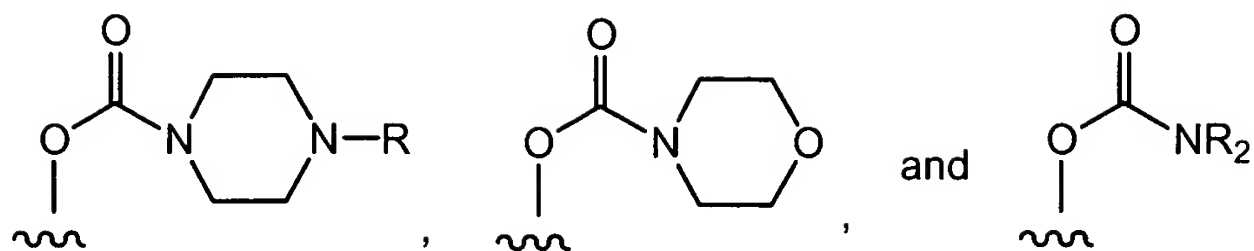
with the proviso that when R¹ is OH, and R², R³, and R⁴ are H, then L is not a bond.

26. (Previously presented): A compound of claim 1 wherein

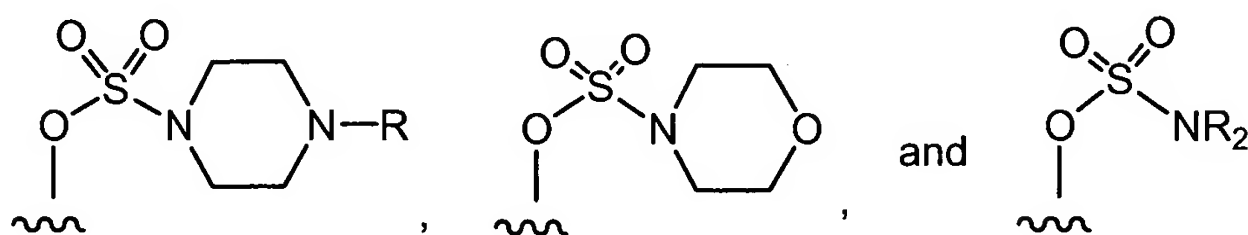
R¹ is selected from CR₃, C(=O)NR₂, OC(=O)OR, OC(=O)NR₂, OC(=O)R, OSO₂NR₂ (sulfamate), NR₂, NRSO₂R, SR, S(O)R, SO₂R and SO₂NR₂ (sulfonamide).

27. (Previously presented): The compound of claim 26 wherein at least one R is a prodrug moiety.

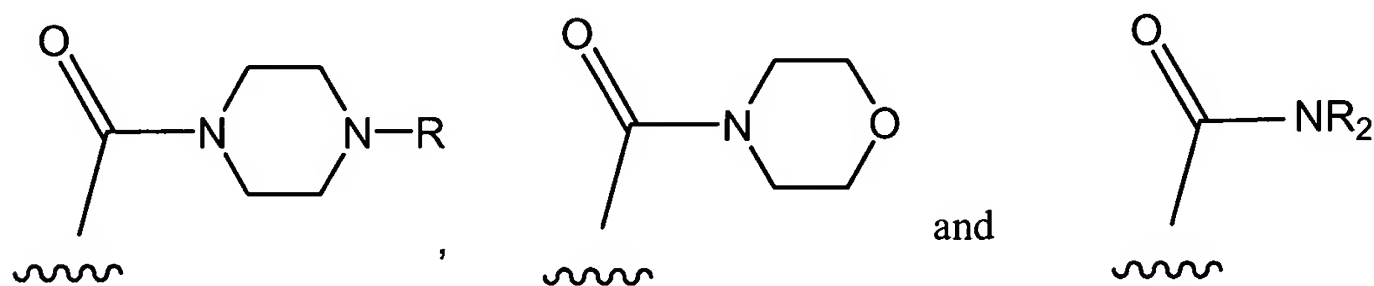
28. (Previously presented): A compound of claim 1 wherein at least one of R¹, R², R³, and R⁴ is selected from the structures:



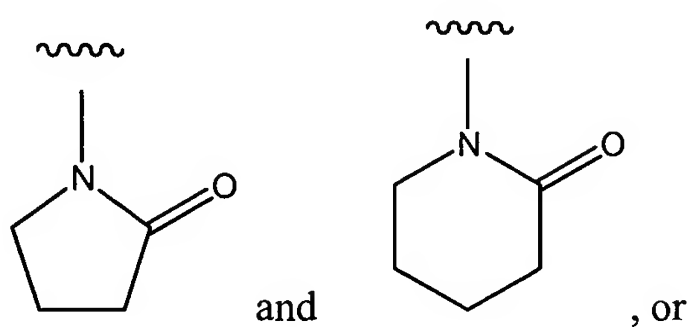
29. (Previously presented): A compound of claim 1 wherein at least one of R^1 , R^2 , R^3 , and R^4 is selected from the structures:



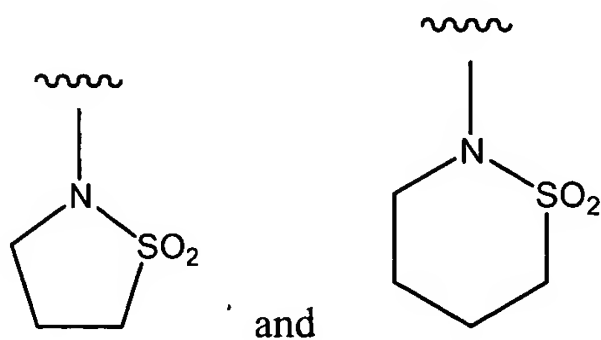
30. (Previously presented): A compound of claim 1 wherein at least one of R^1 , R^2 , R^3 , and R^4 is selected from the structures:



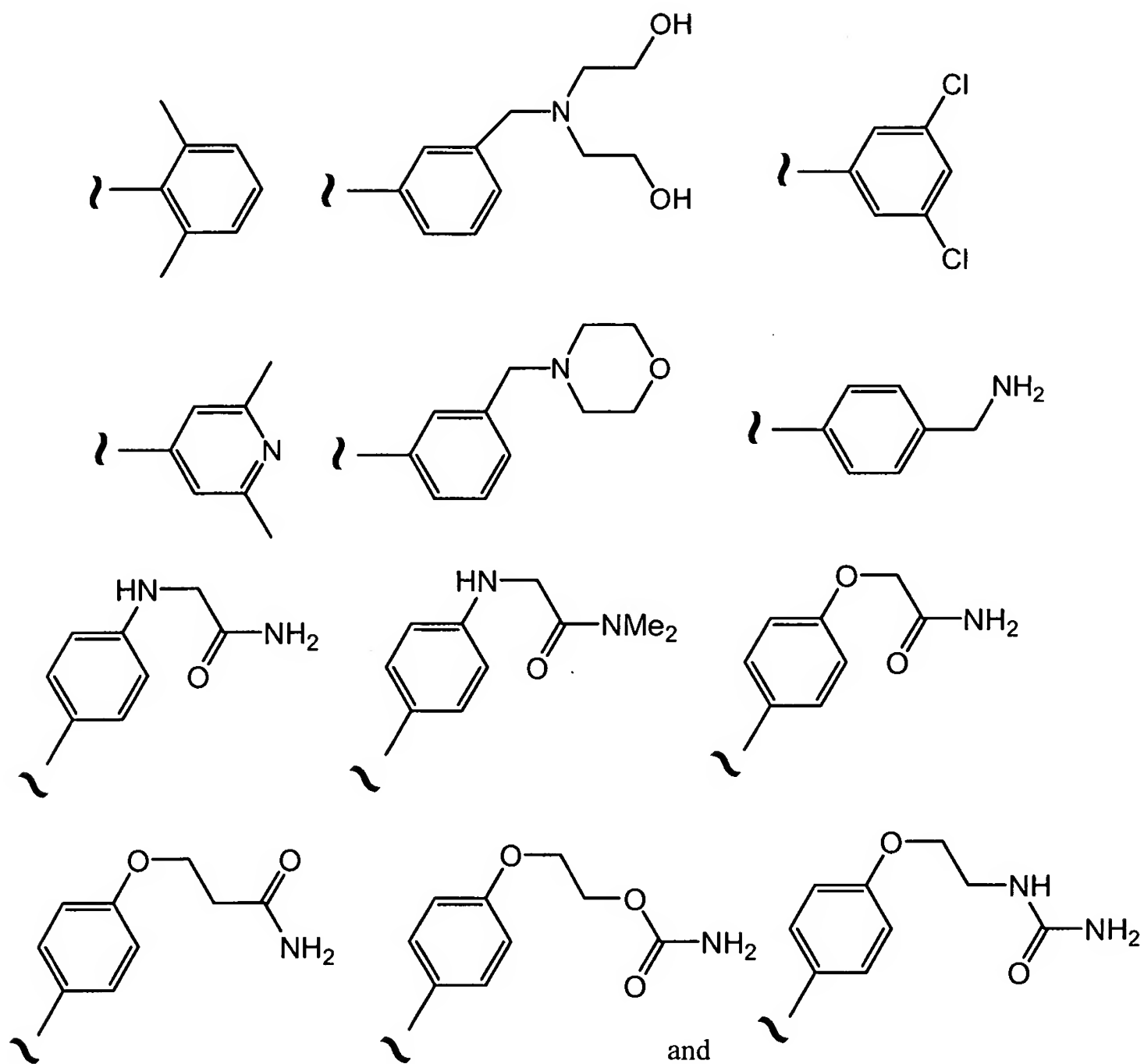
31. (Previously presented): A compound of claim 1 wherein at least one of R^1 , R^2 , R^3 , and R^4 is a lactam having the structures:



a sultam having the structures:

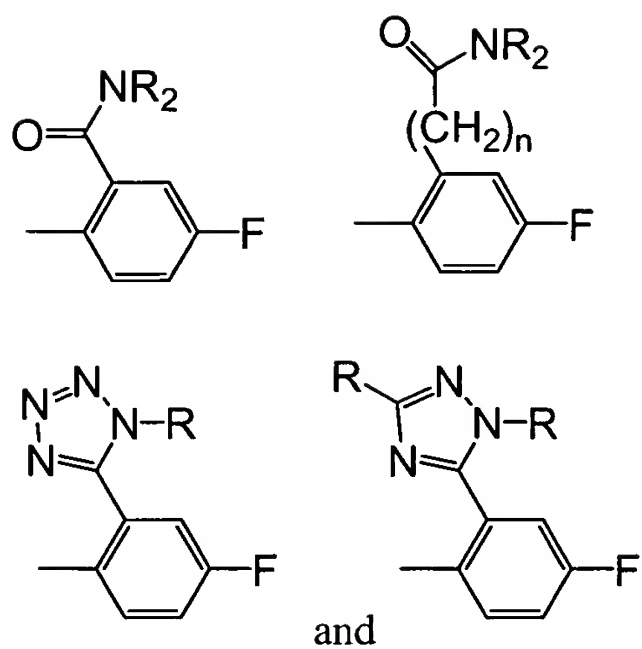


32. (Previously presented): A compound of claim 1 wherein Ar is selected from the structures:



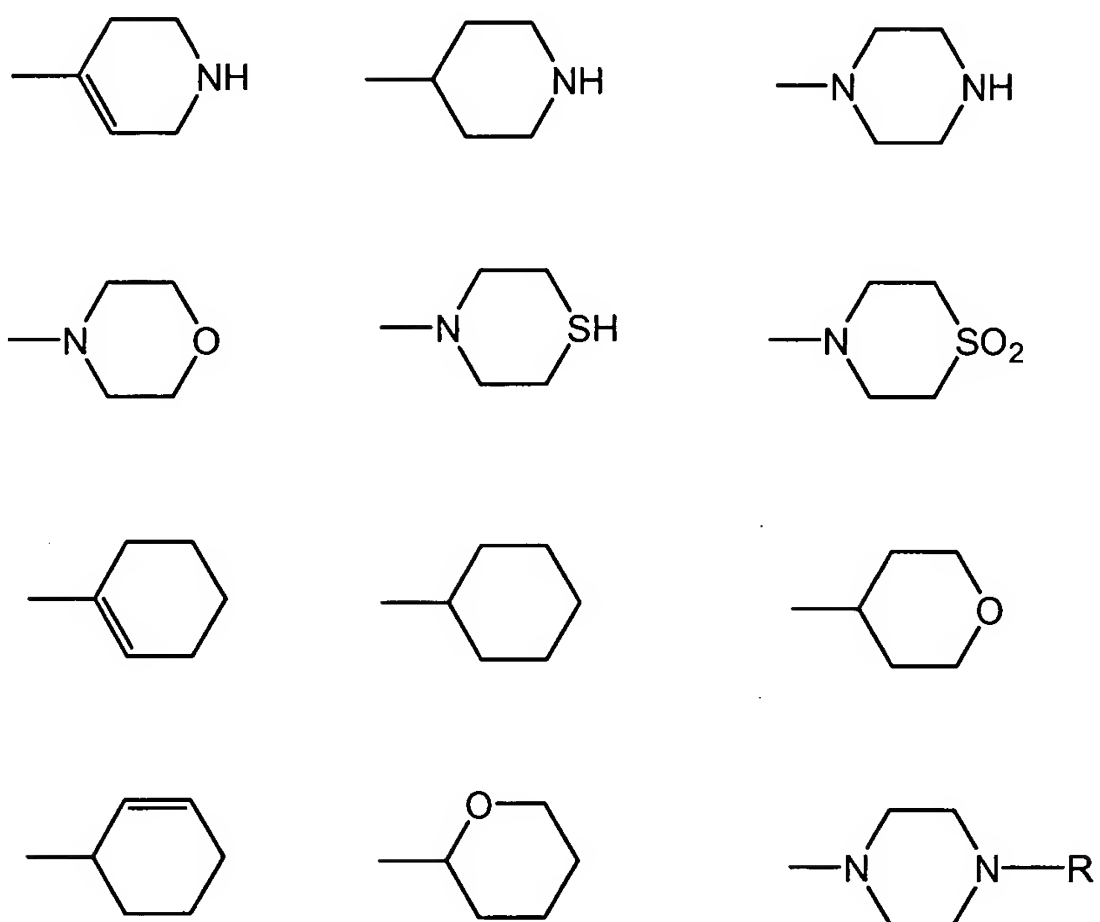
where the wavy line indicates the covalent attachment site to L.

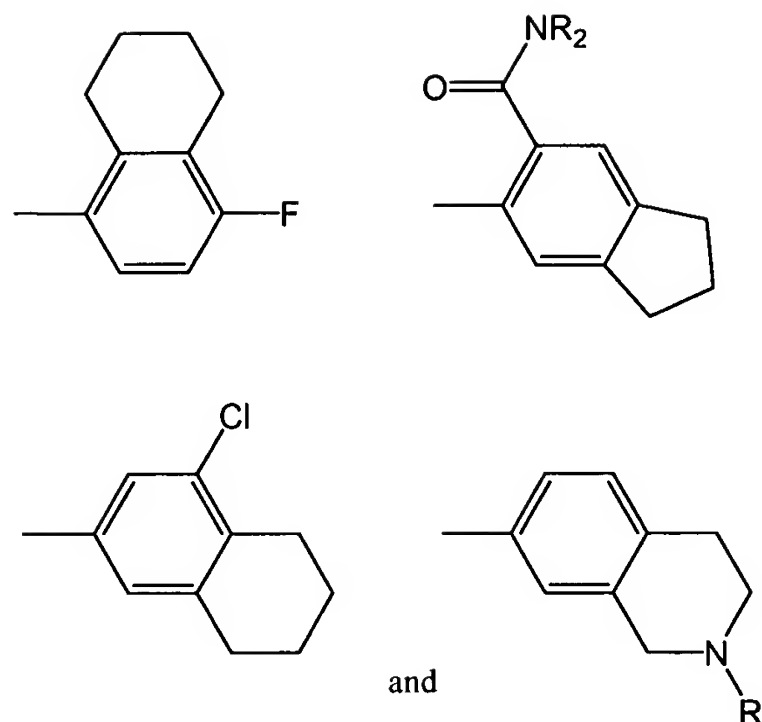
33. (Previously presented): A compound of claim 1 wherein Ar is selected from the structures:



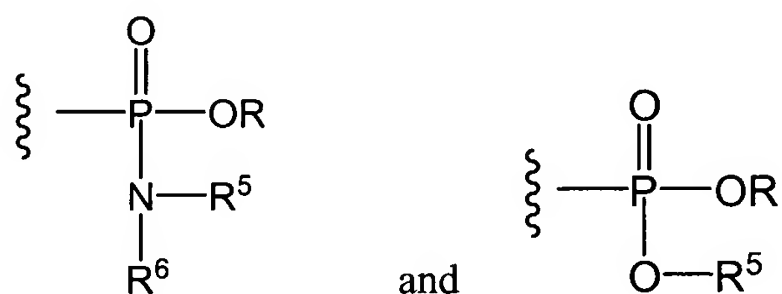
where n is 1 to 6.

34. (Previously presented): A compound of claim 1 wherein Ar is selected from the structures:



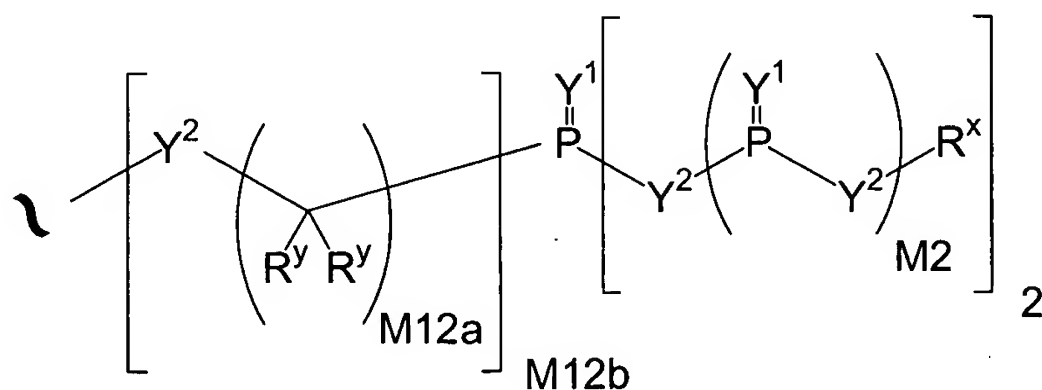


35. (Previously presented): A compound of claim 1 comprising a prodrug moiety selected from the structures:



wherein R^5 is $-\text{CR}_2\text{CO}_2R^7$ where R^6 and R^7 are independently H or $\text{C}_1\text{--C}_8$ alkyl.

36. (Previously presented): The compound of claim 1 comprising a phosphonate or prodrug moiety having the structure:



wherein:

Y^1 is independently O, S, $\text{N}(\text{R}^x)$, $\text{N}(\text{O})(\text{R}^x)$, $\text{N}(\text{OR}^x)$, $\text{N}(\text{O})(\text{OR}^x)$, or $\text{N}(\text{N}(\text{R}^x)_2)$;

Y^2 is independently a bond, O, $\text{N}(\text{R}^x)$, $\text{N}(\text{O})(\text{R}^x)$, $\text{N}(\text{OR}^x)$, $\text{N}(\text{O})(\text{OR}^x)$, $\text{N}(\text{N}(\text{R}^x)_2)$, $-\text{S}(\text{O})-$ (sulfoxide), $-\text{S}(\text{O})_2-$ (sulfone), $-\text{S}-$ (sulfide), or $-\text{S}-\text{S}-$ (disulfide);

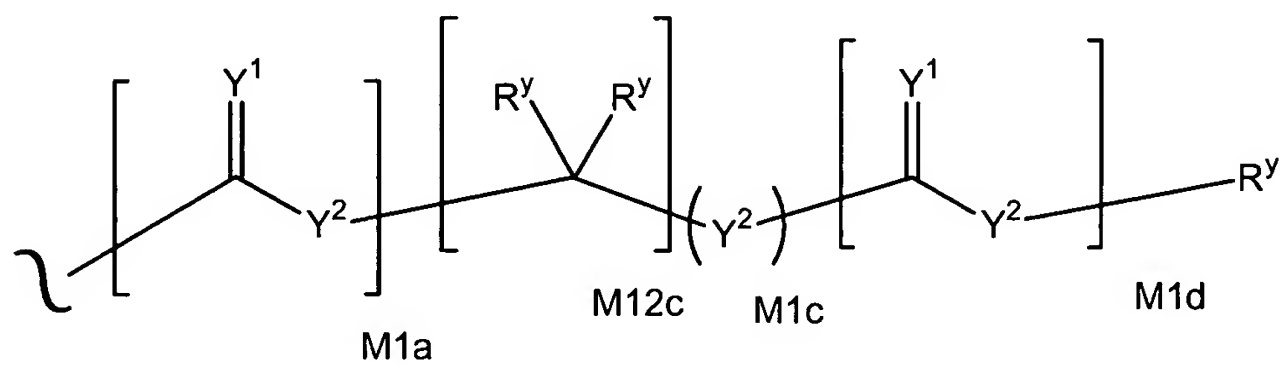
M2 is 0, 1 or 2;

M12a is 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 11, or 12;

M12b is 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 11, or 12;

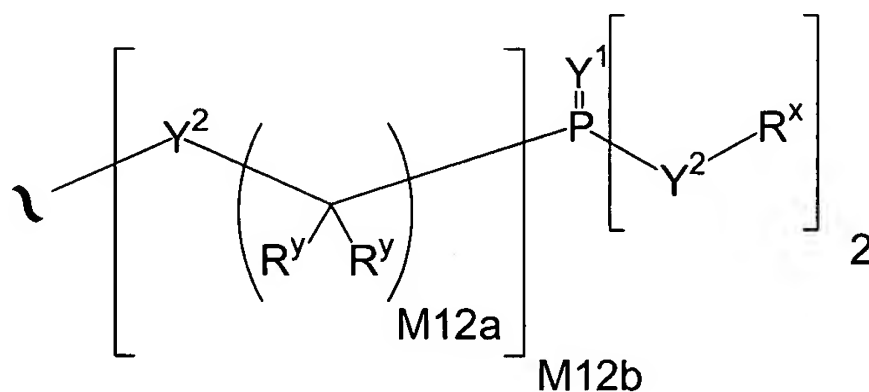
R^y is independently H, C_1-C_6 alkyl, C_1-C_6 substituted alkyl, C_6-C_{20} aryl, C_6-C_{20} substituted aryl, or a protecting group, or where taken together at a carbon atom, two vicinal R^y groups form a carbocycle or a heterocycle; and

R^x is independently H, C_1-C_6 alkyl, C_1-C_6 substituted alkyl, C_6-C_{20} aryl, C_6-C_{20} substituted aryl, or a protecting group, or the formula:

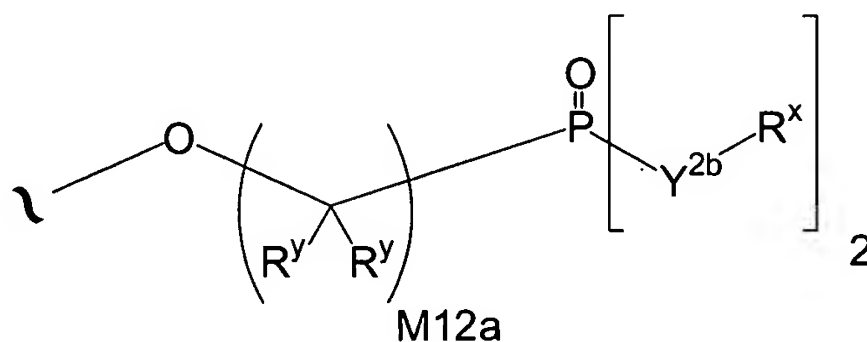


where M1a, M1c, and M1d are independently 0 or 1, and M12c is 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 11 or 12.

37. (Previously presented): The compound of claim 36 wherein the phosphonate or prodrug moiety has the structure:

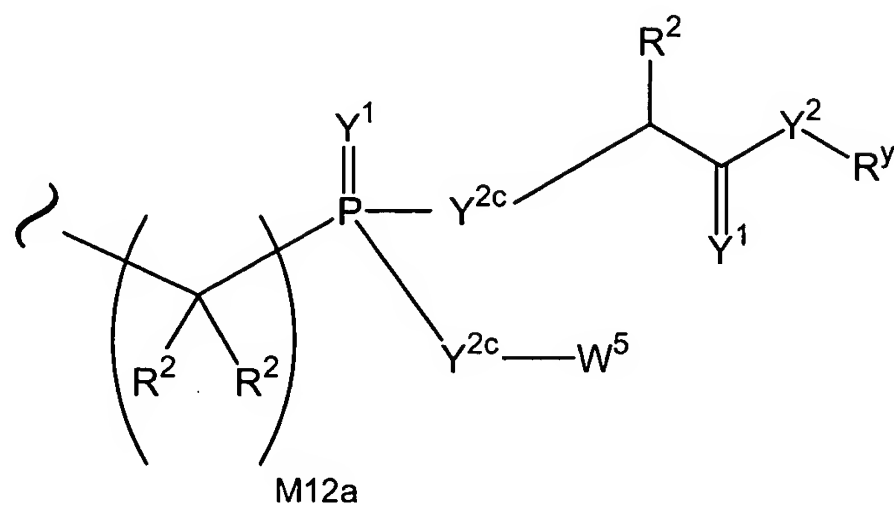


38. (Previously presented): The compound of claim 37 wherein the phosphonate or prodrug moiety has the structure:



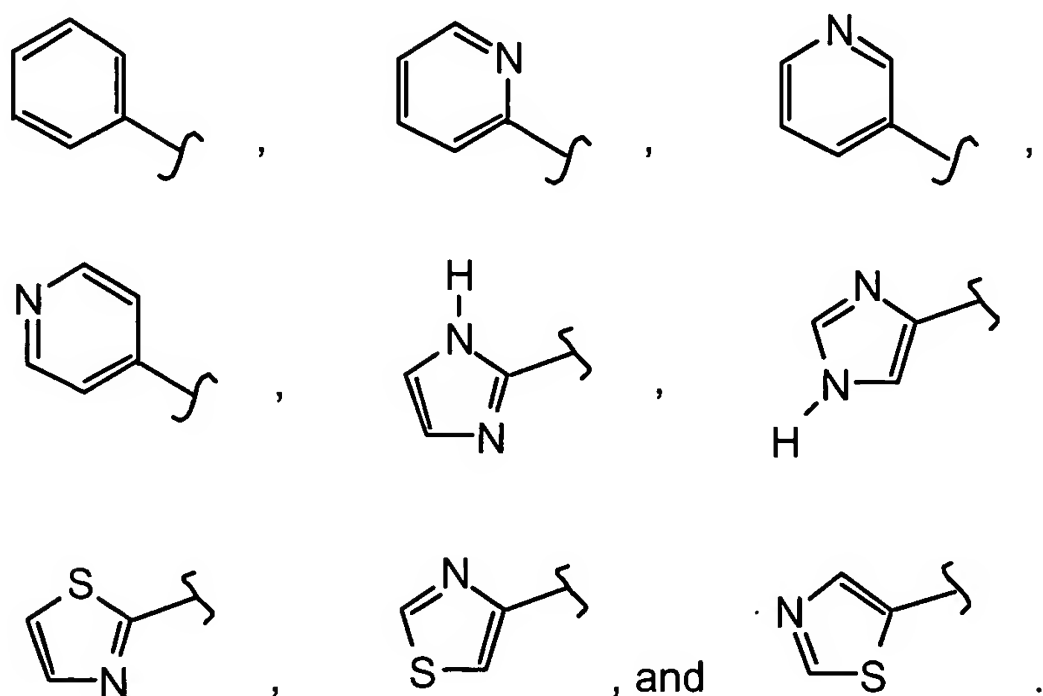
where Y^{2b} is O or $N(R^x)$.

39. (Previously presented): The compound of claim 37 wherein the phosphonate or prodrug moiety has the structure:

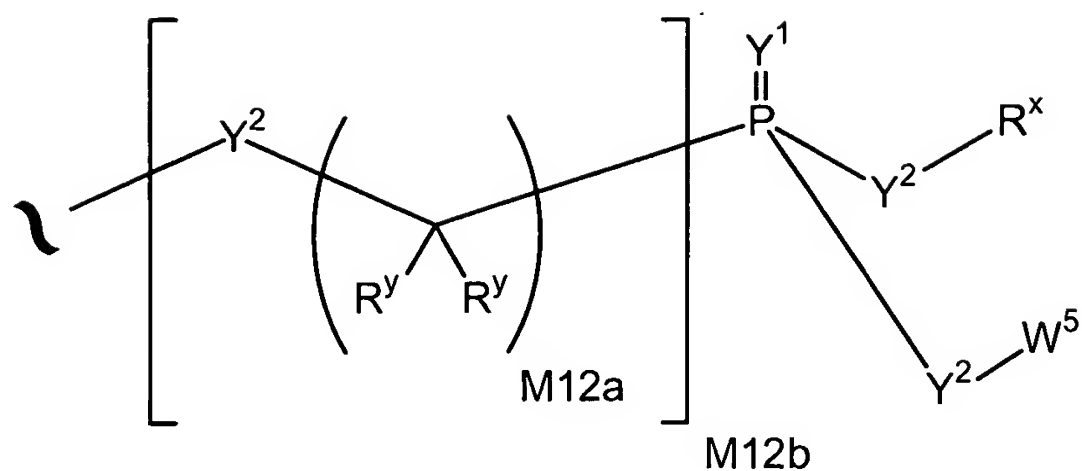


where W^5 is a carbocycle, and Y^{2c} is O, $N(R^y)$ or S.

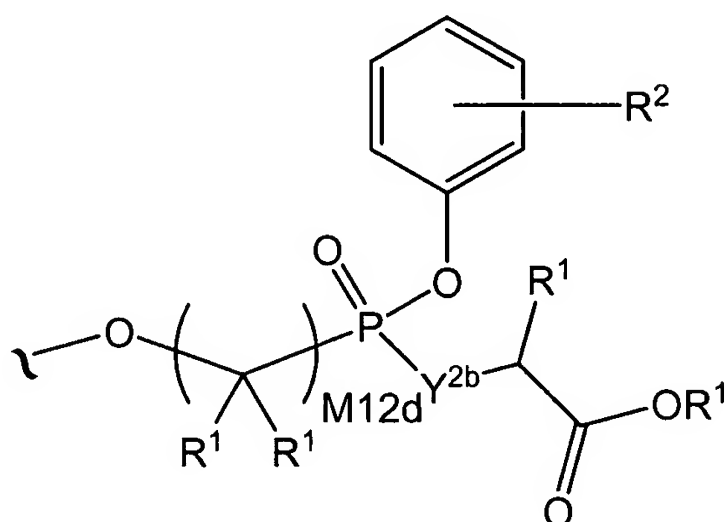
40. (Previously presented): The compound of claim 39 wherein W^5 is selected from the structures:



41. (Previously presented): The compound of claim 37 wherein the phosphonate or prodrug moiety has the structure:

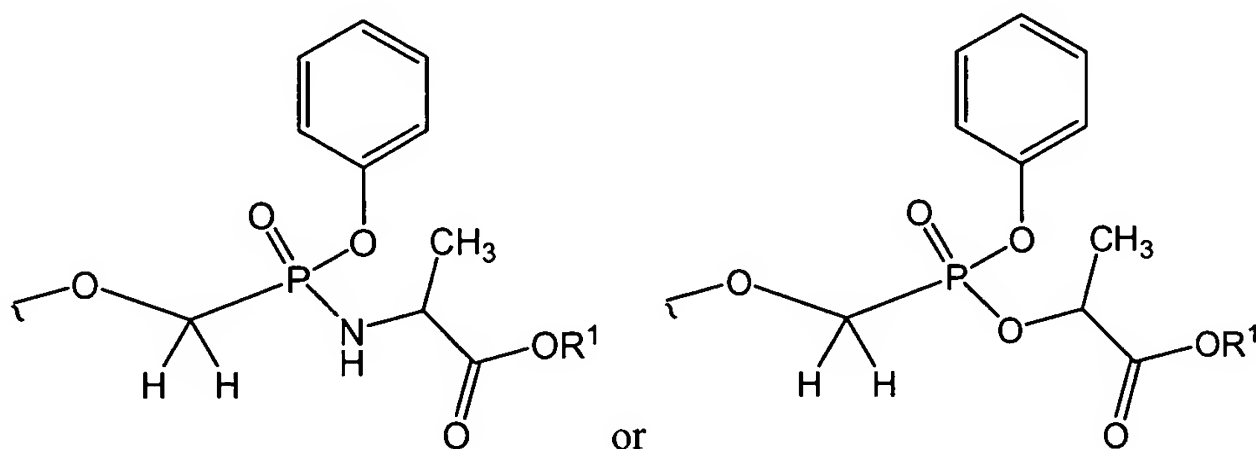


42. (Previously presented): The compound of claim 1 wherein the phosphonate or prodrug moiety has the structure:

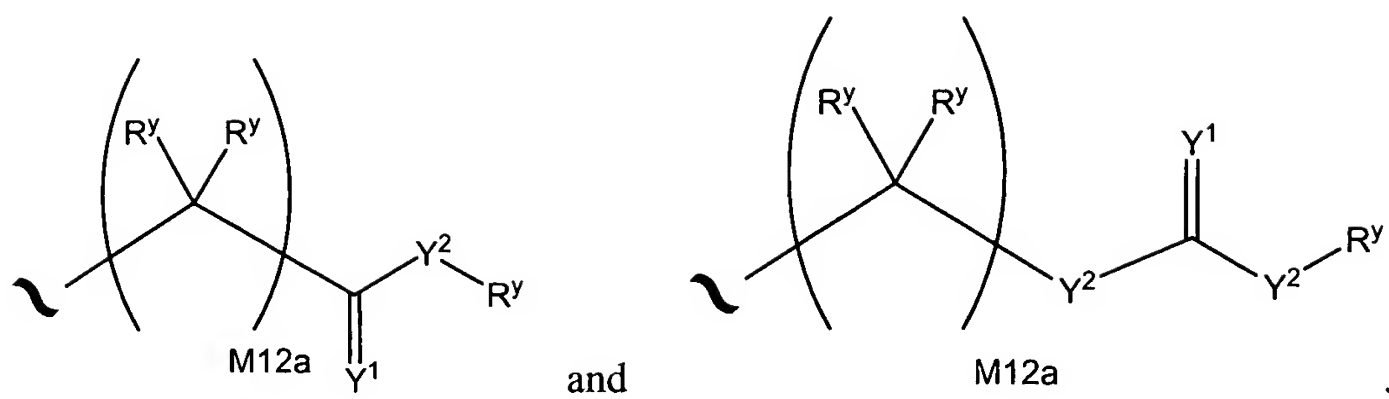


wherein Y^{2b} is O or $N(R^x)$; M12d is 1, 2, 3, 4, 5, 6, 7 or 8; R^1 is H or C_1-C_6 alkyl; and the phenyl carbocycle is substituted with 0 to 3 R^2 groups where R^2 is C_1-C_6 alkyl or substituted alkyl.

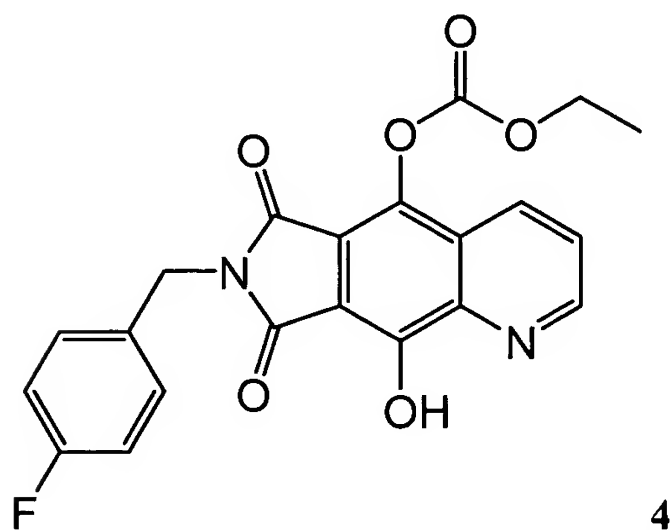
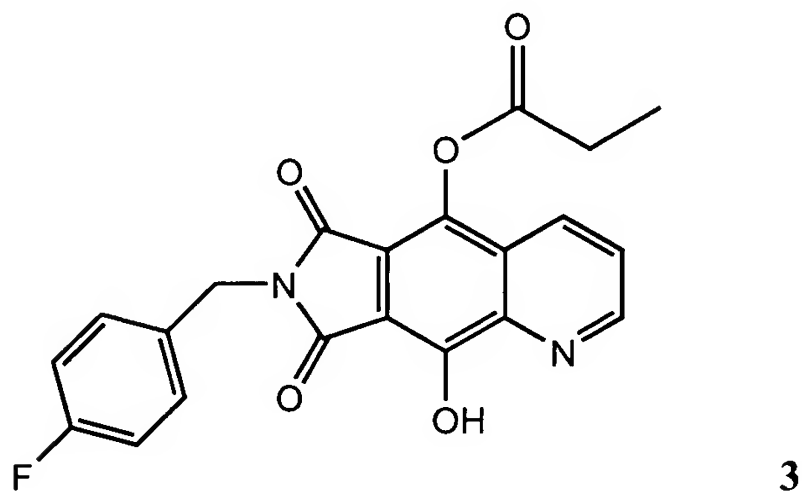
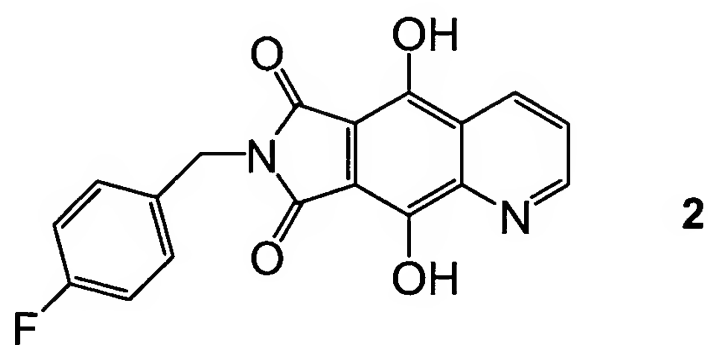
43. (Previously presented): The compound of claim 1 wherein the phosphonate or prodrug moiety has the structure:

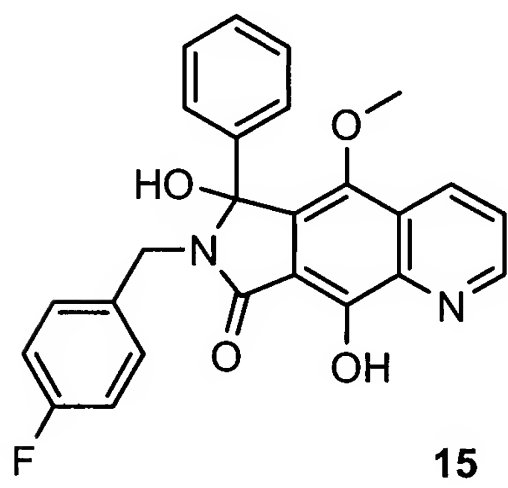
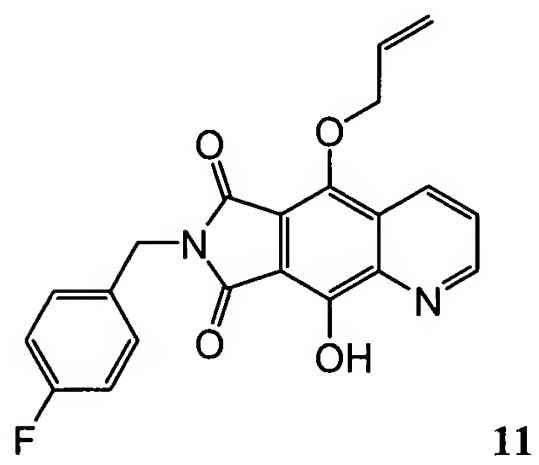
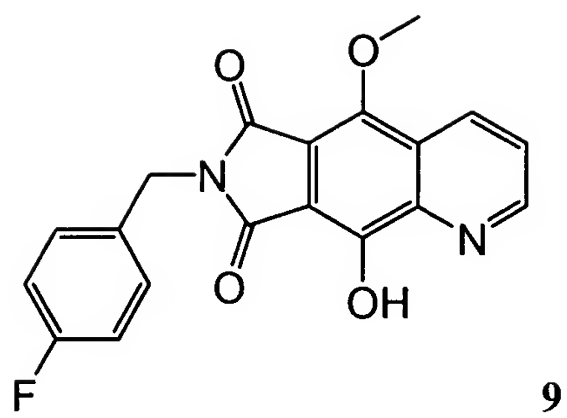


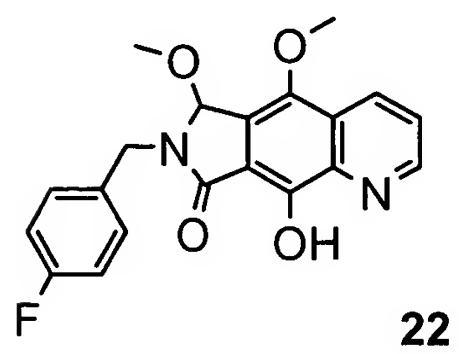
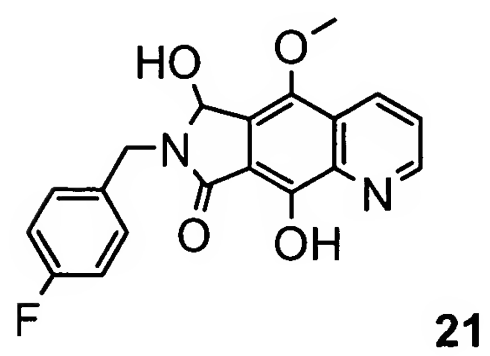
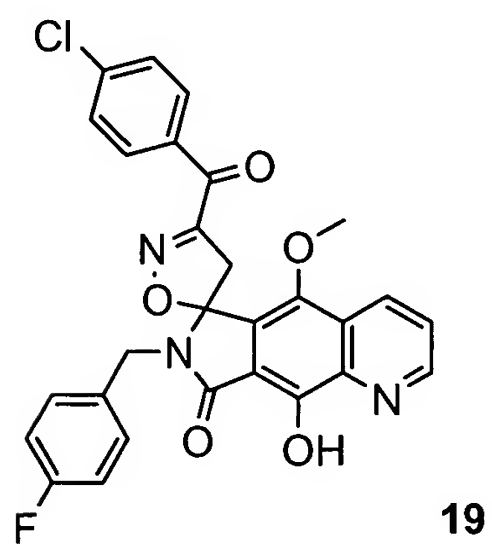
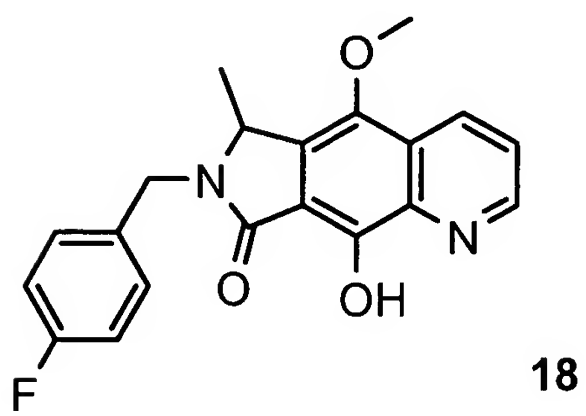
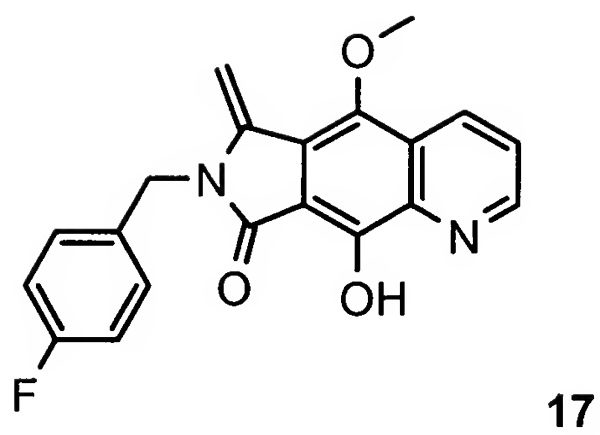
44. (Previously presented): The compound of claim 36 wherein R^x is selected from the structures:

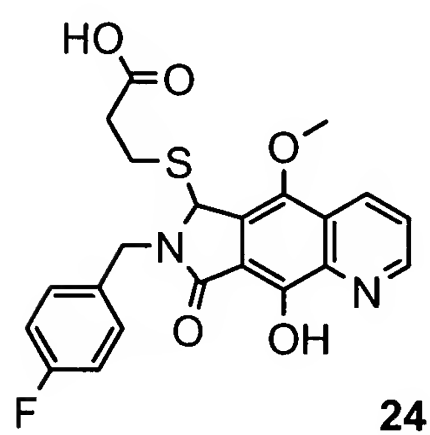
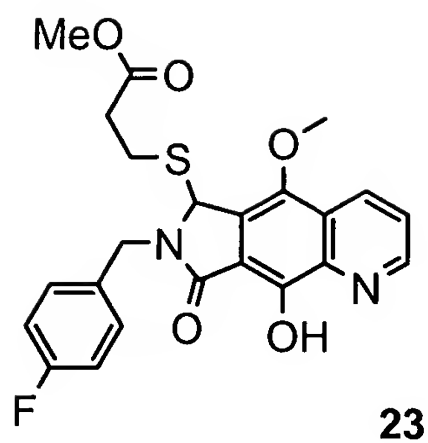


45. (Previously presented): A compound selected from the structures:

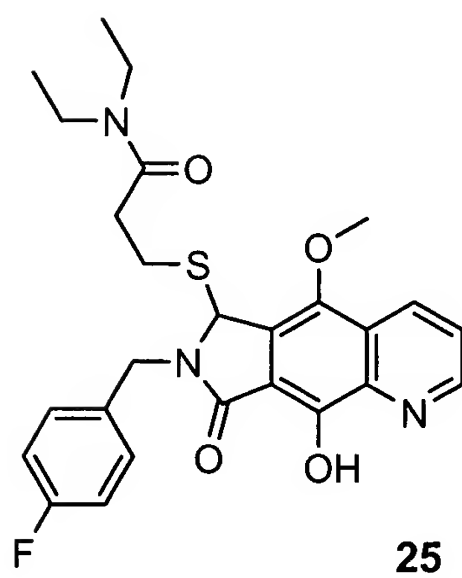






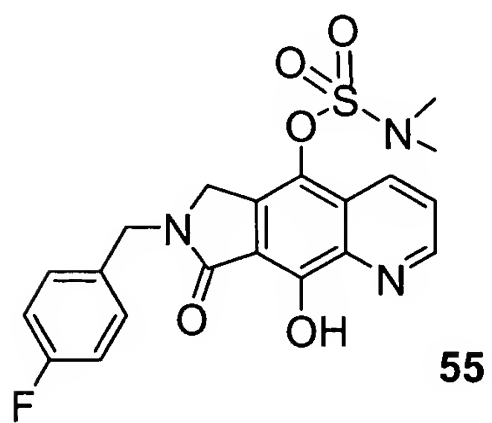
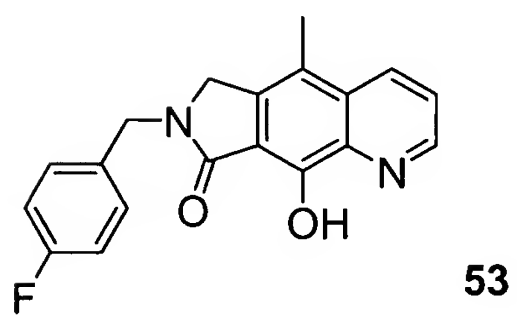
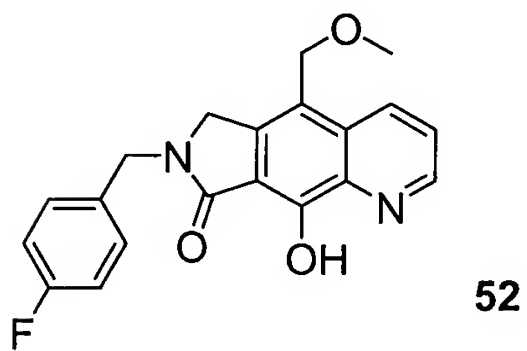
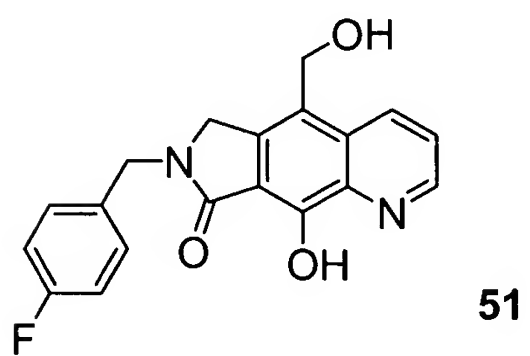
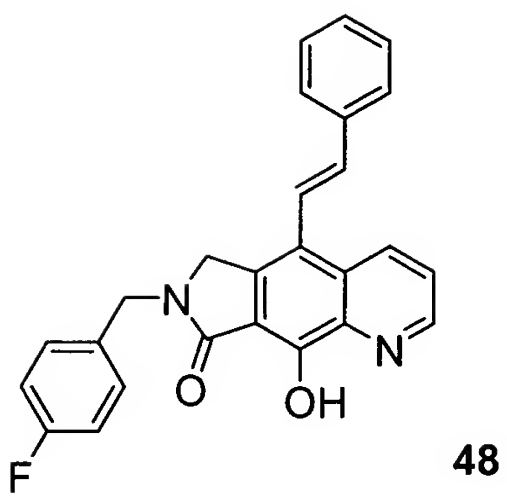


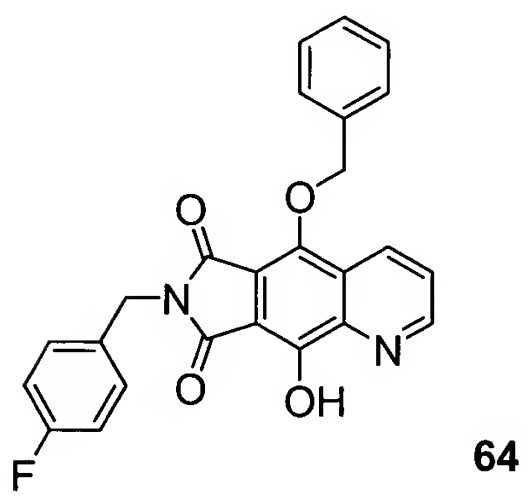
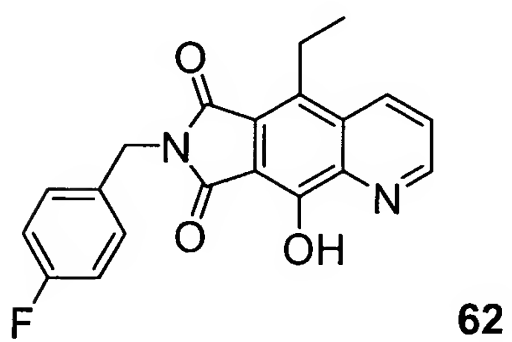
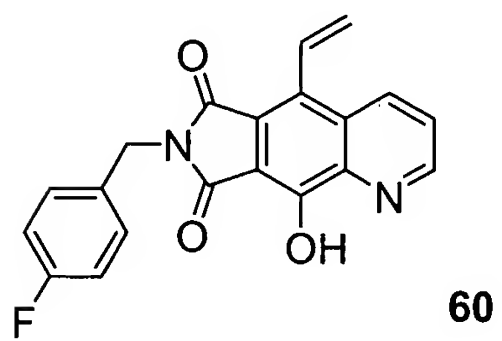
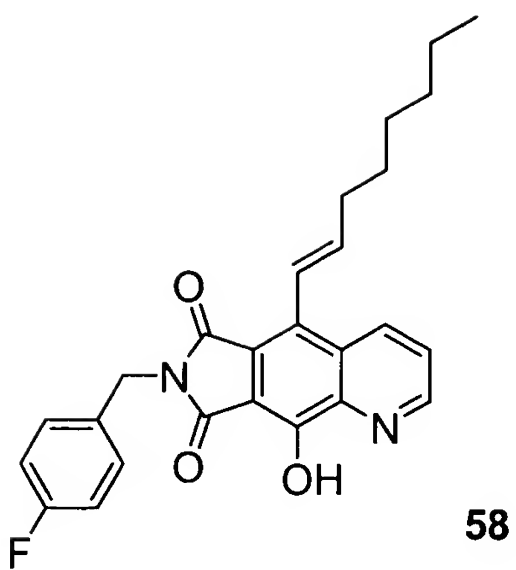
and

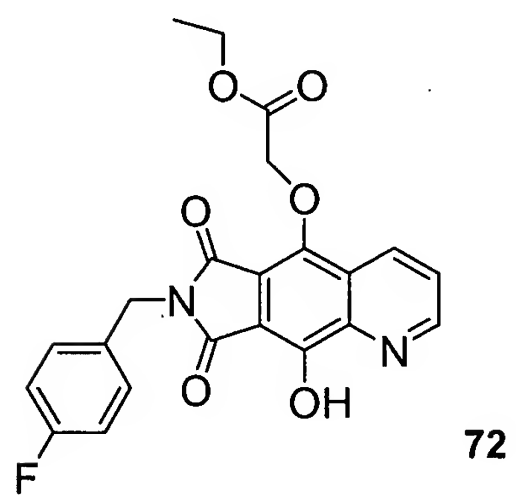
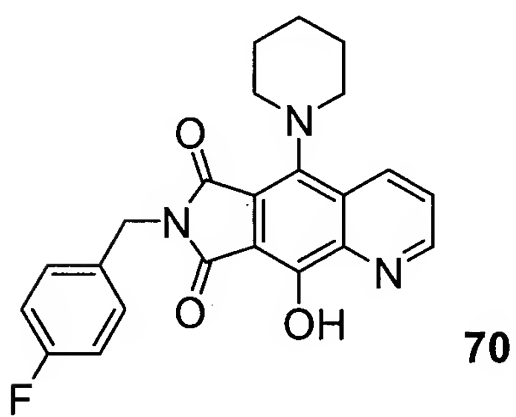
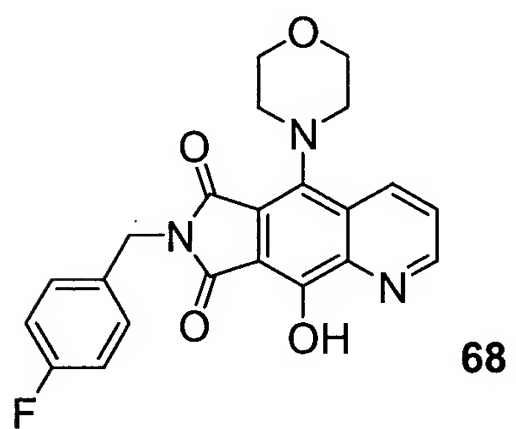
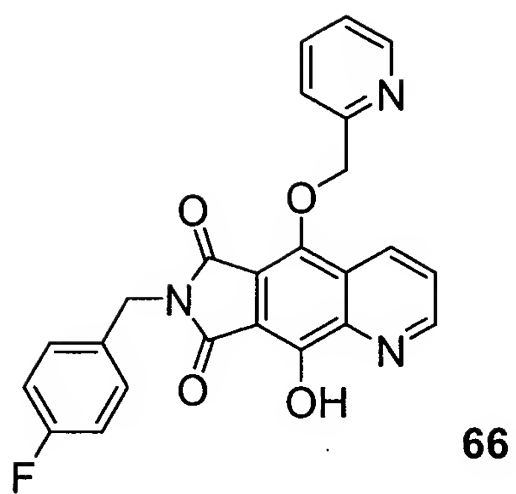


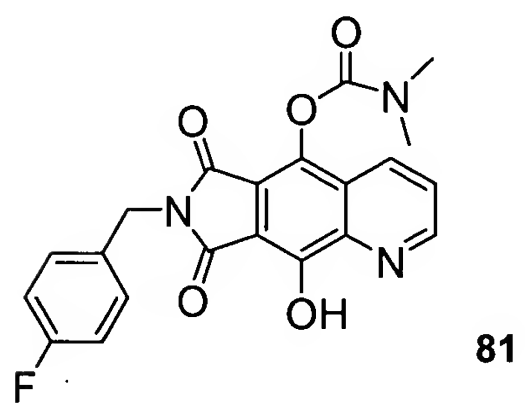
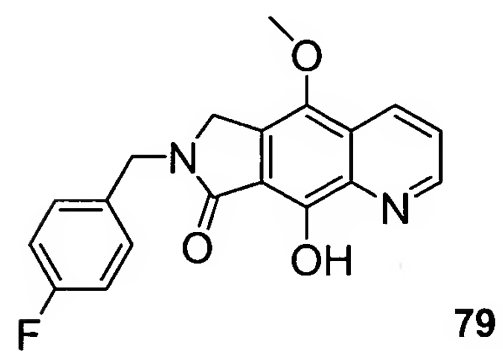
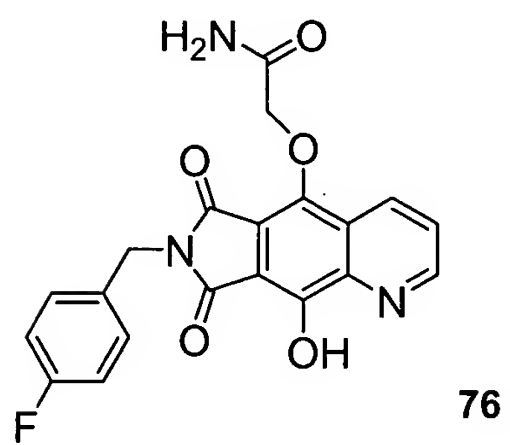
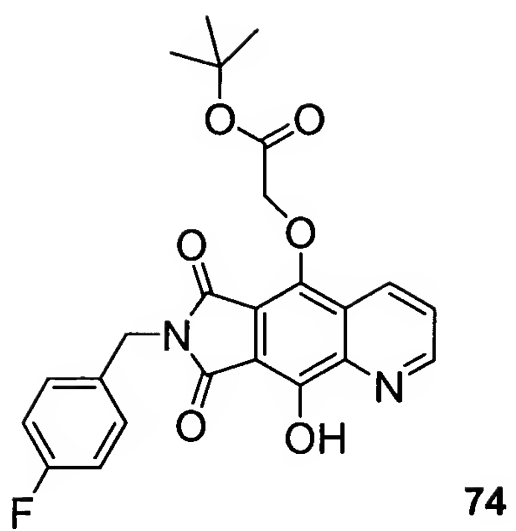
46. (Previously presented): A compound of claim 1 wherein none of R^2 , R^3 , R^4 , R , or R^{x2} is a prodrug moiety.

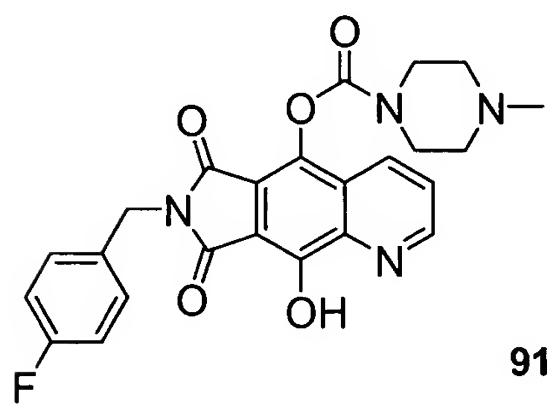
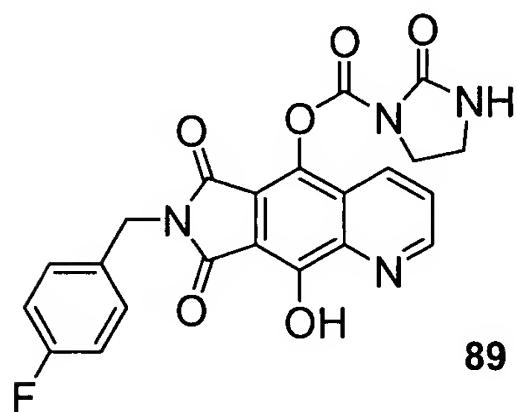
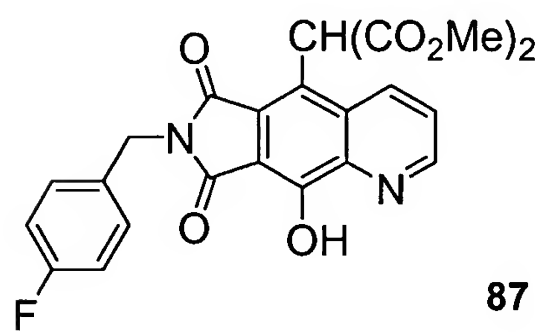
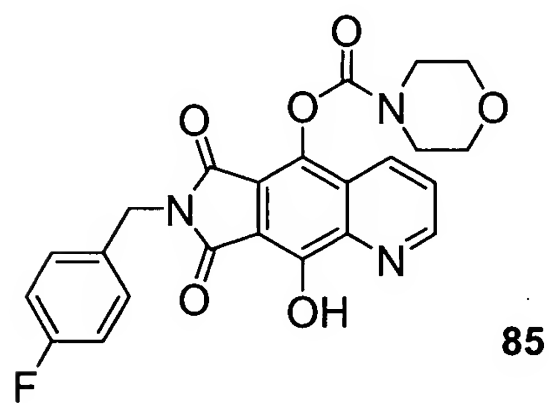
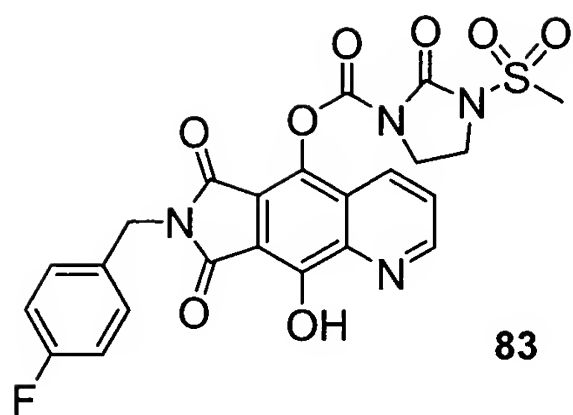
47. (Previously presented): A compound selected from the structures:

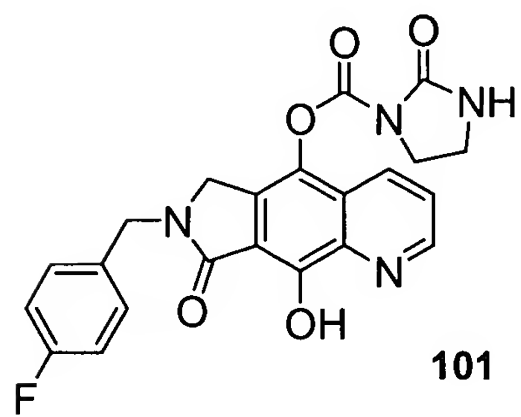
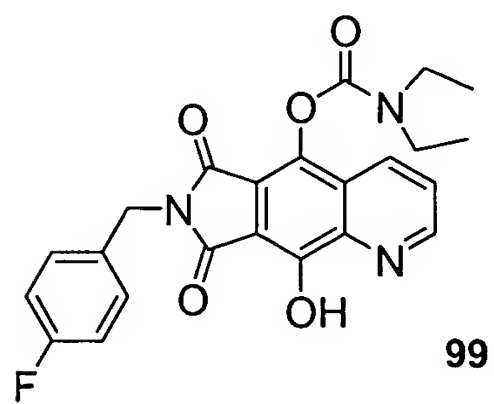
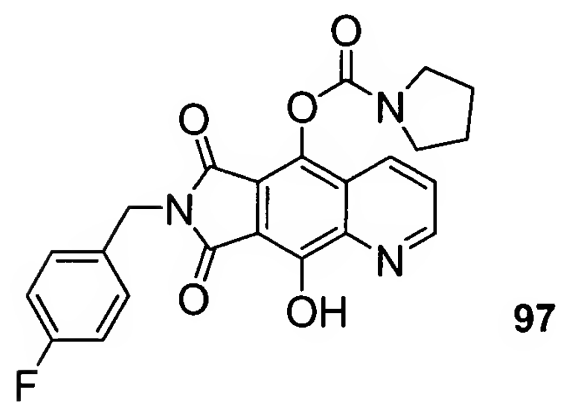
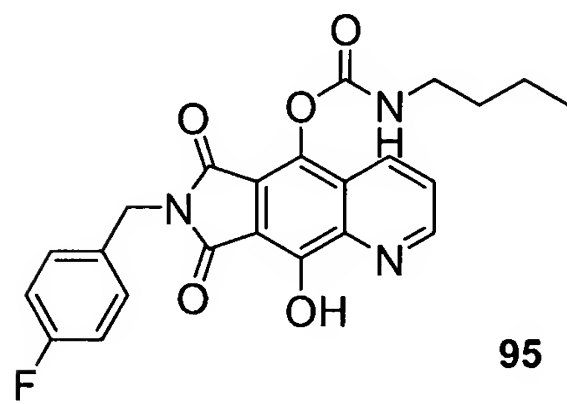
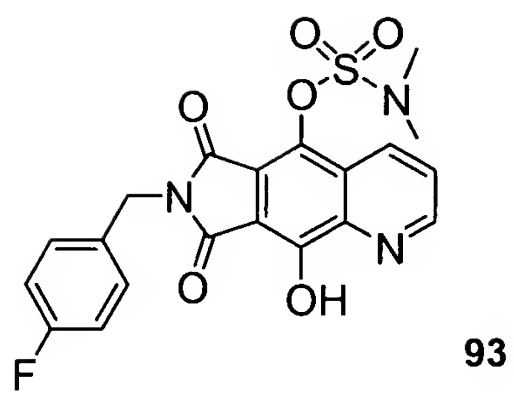


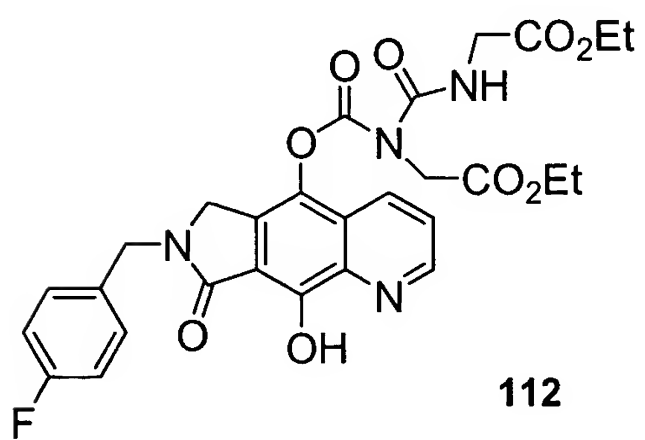
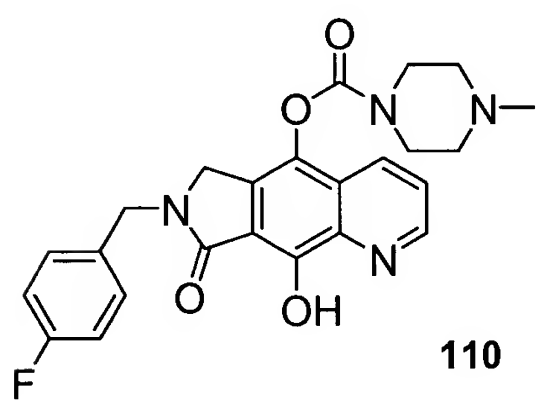
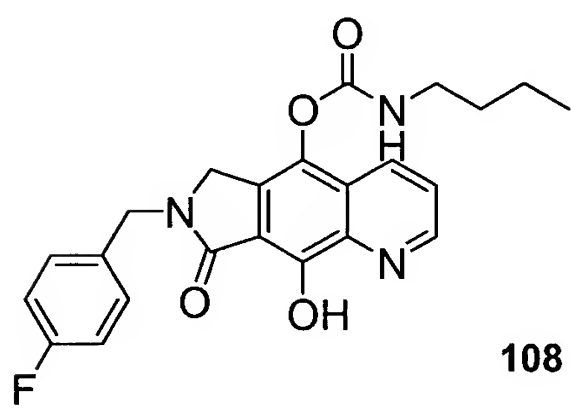
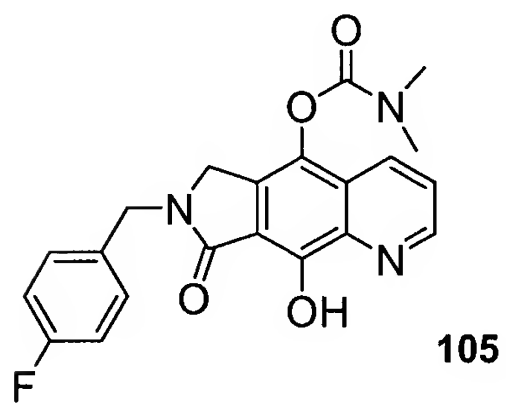
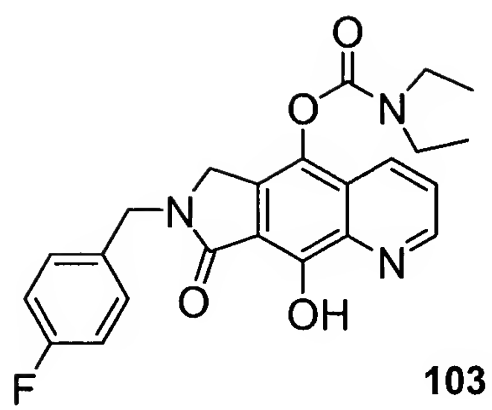


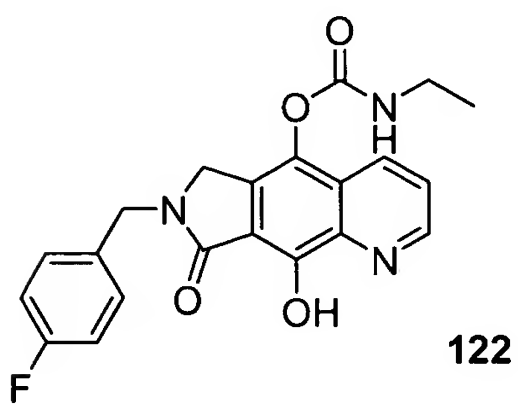
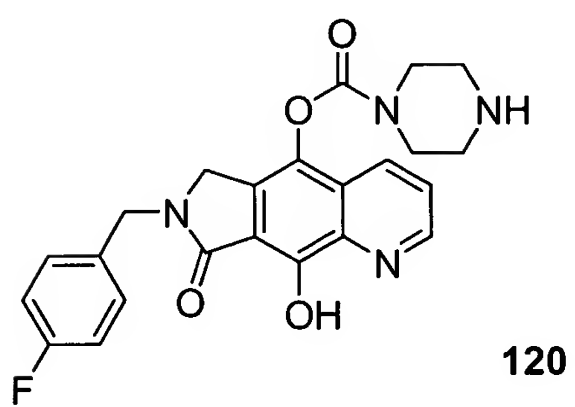
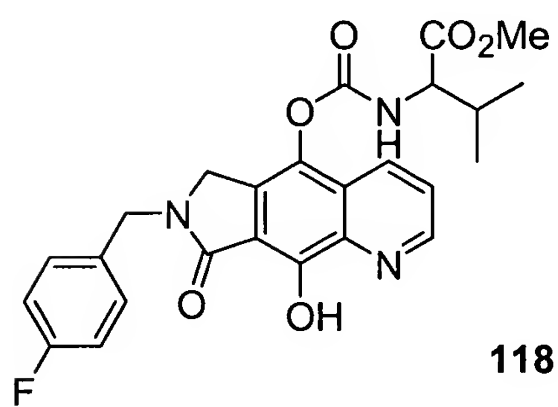
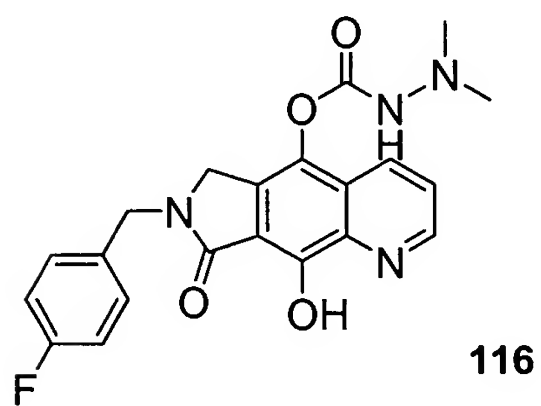
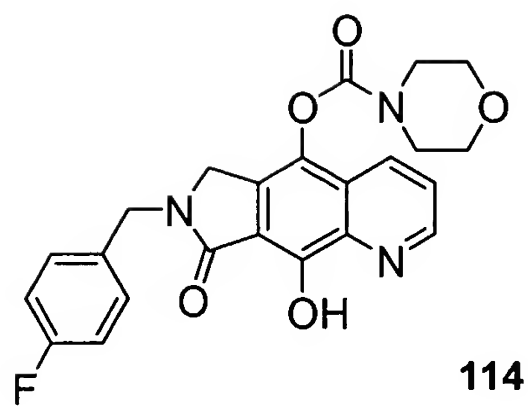


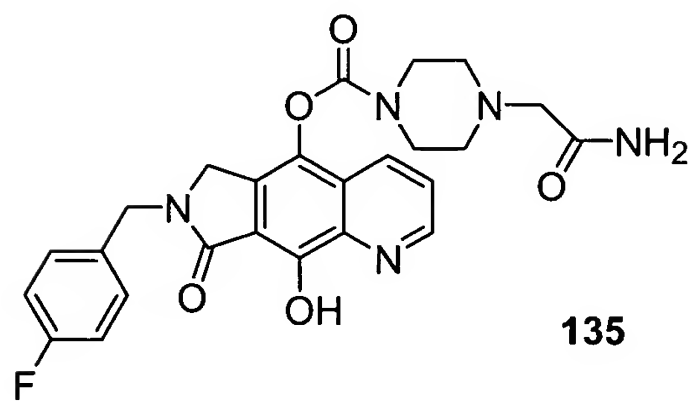
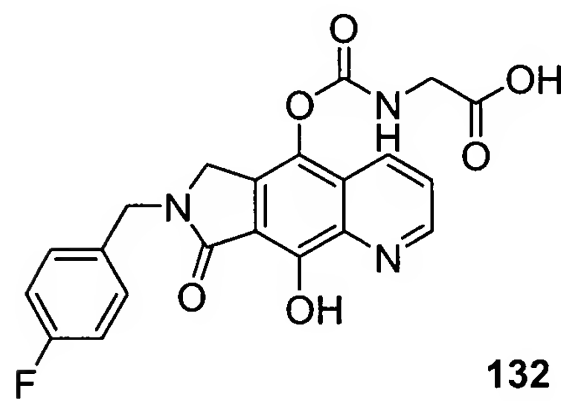
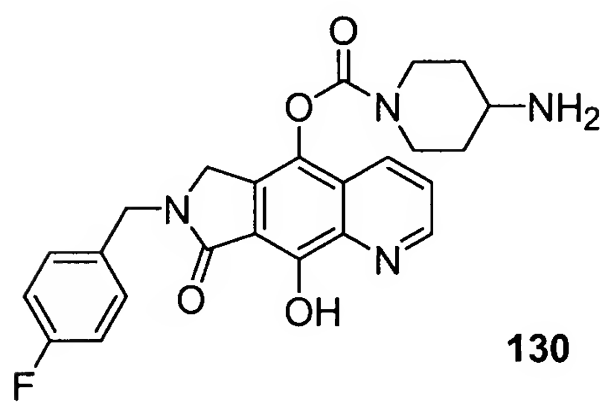
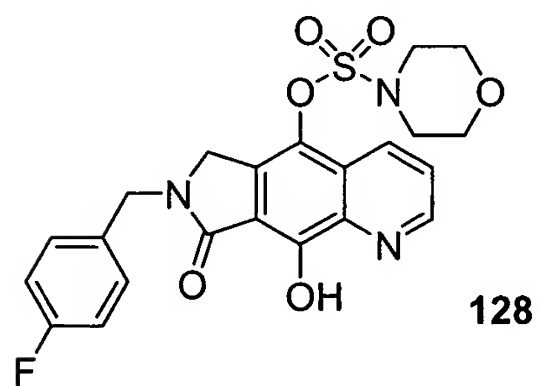
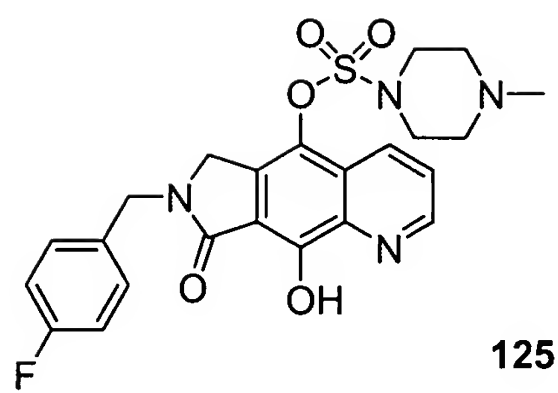


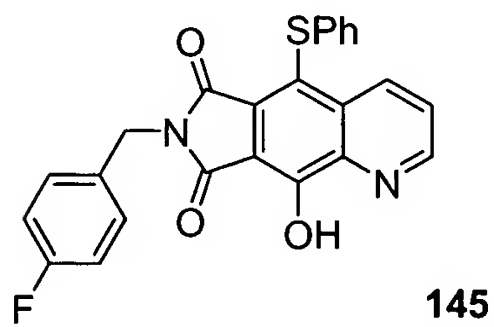
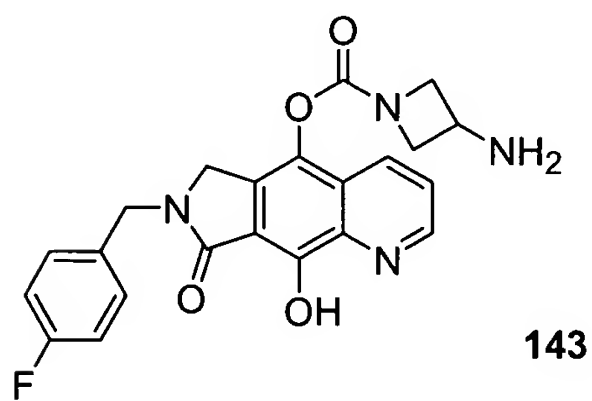
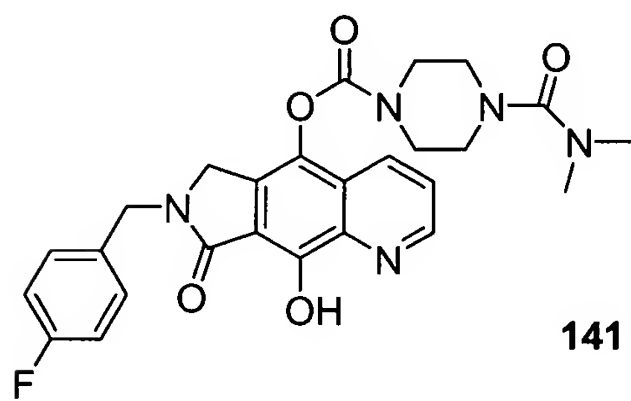
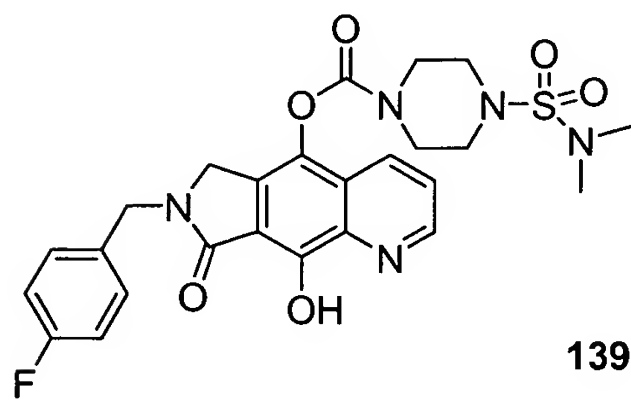
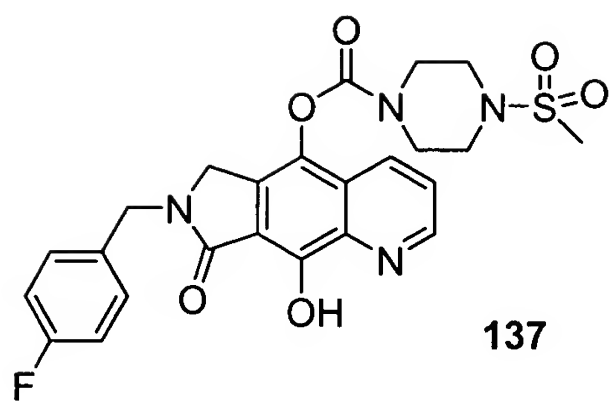


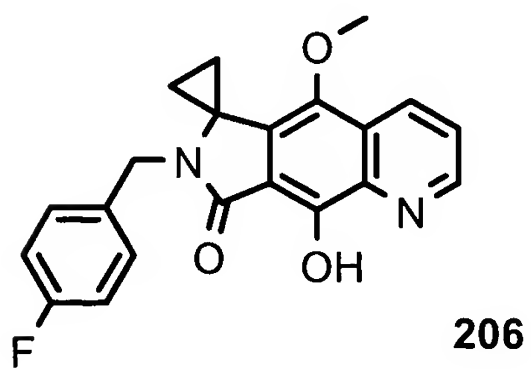
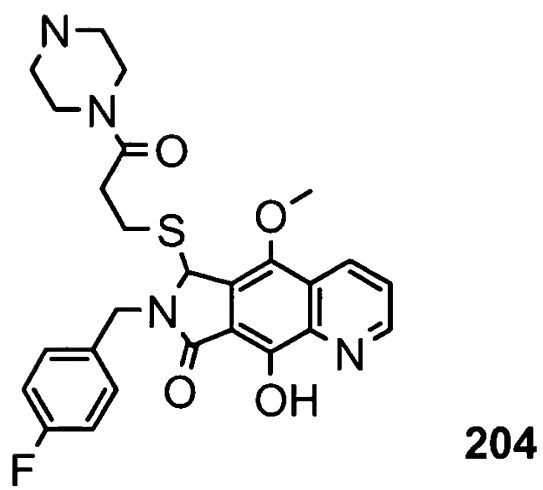
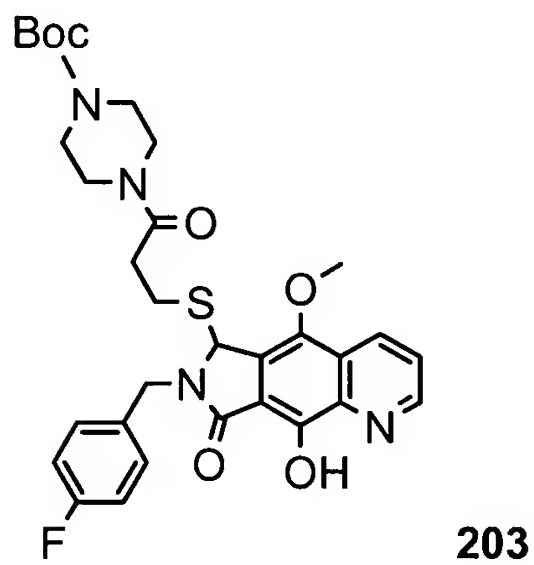
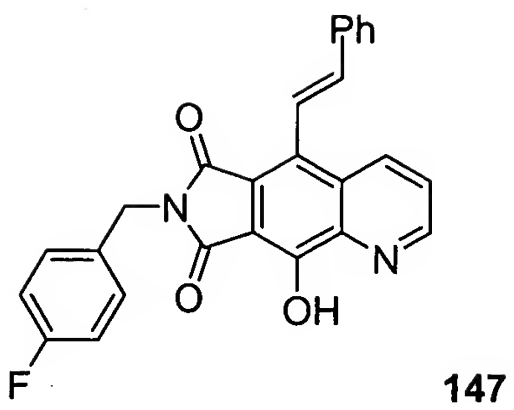


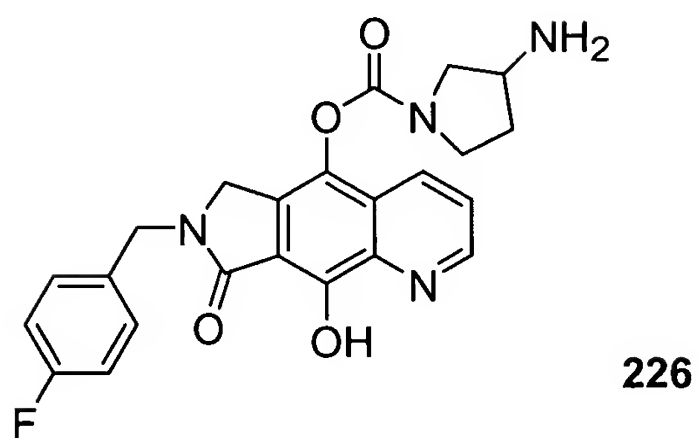
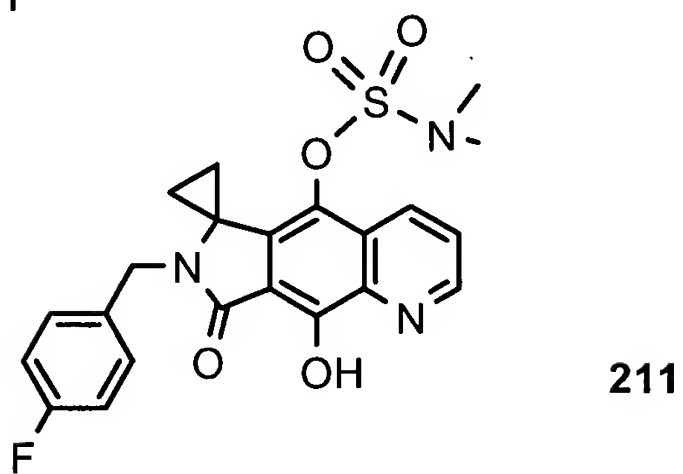
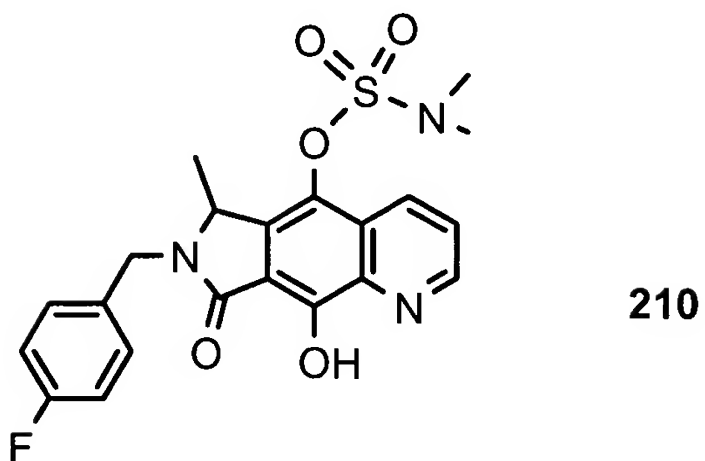
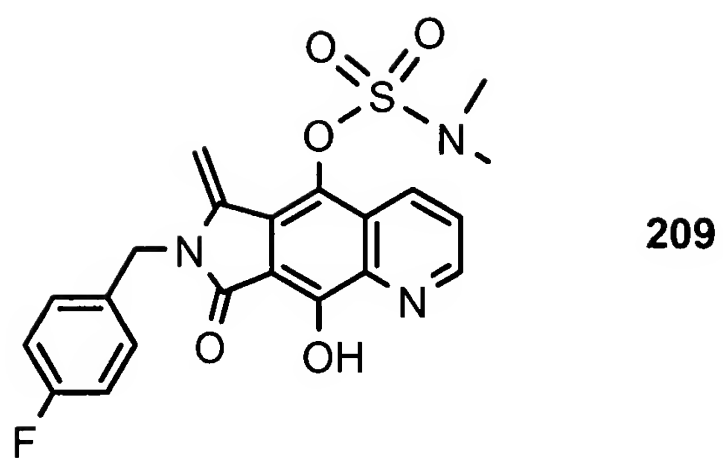


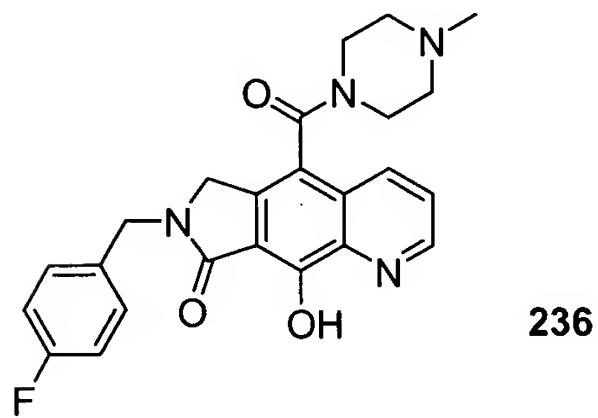
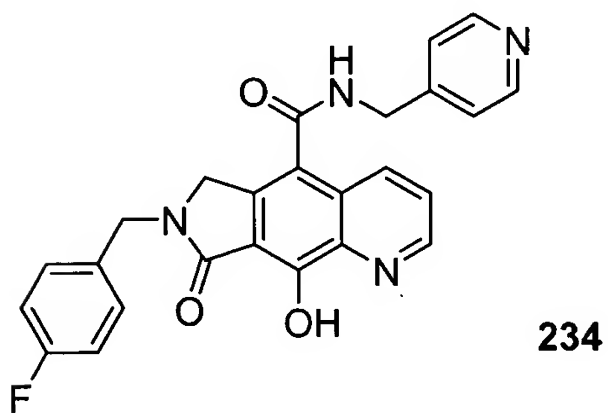
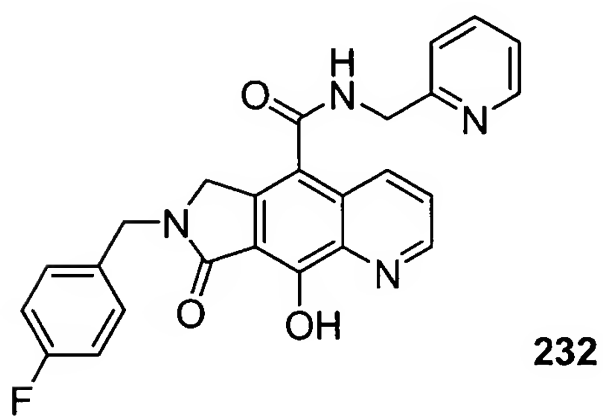
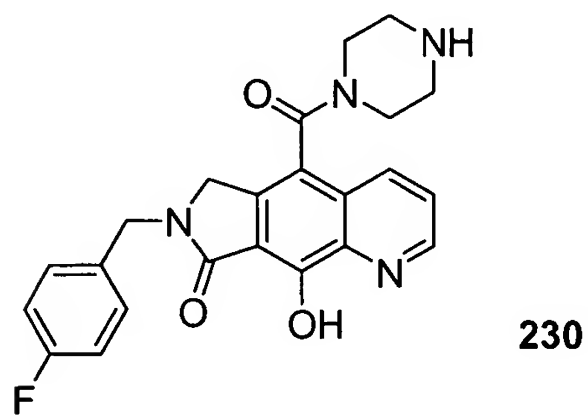
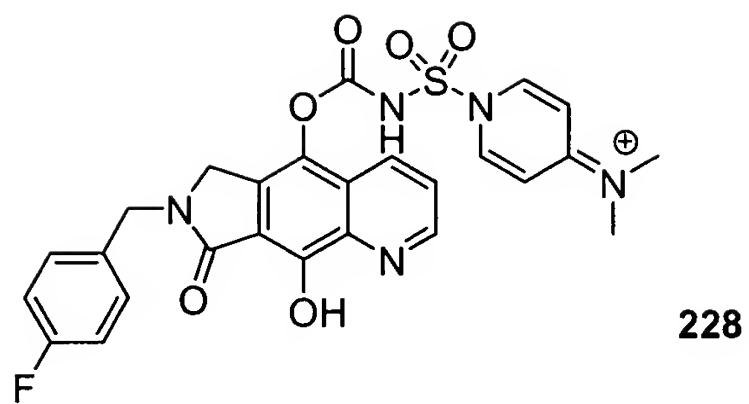


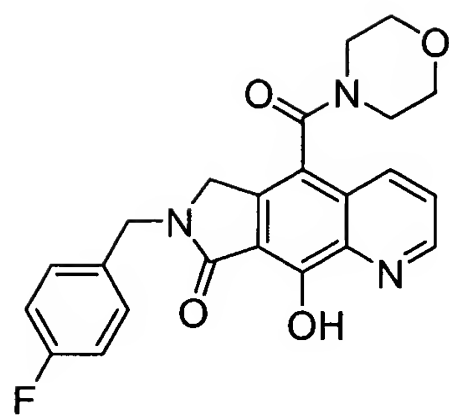




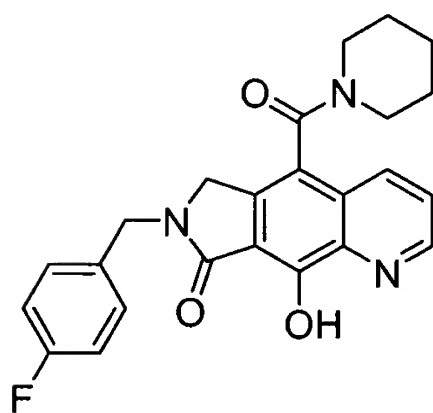




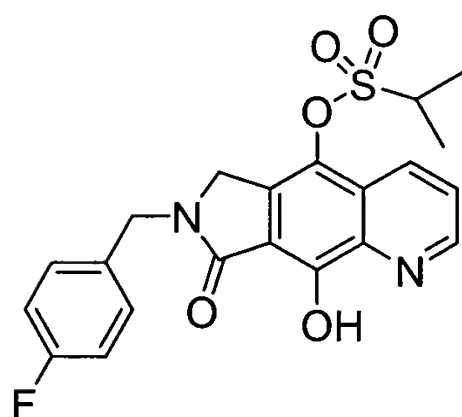




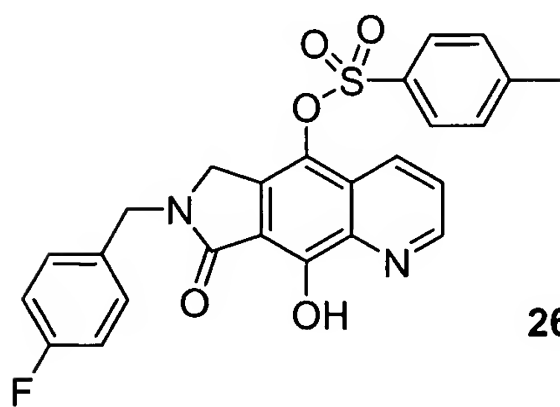
238



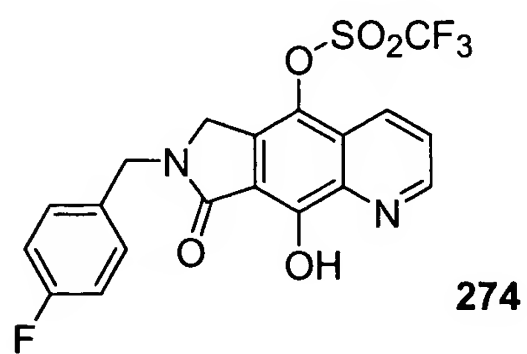
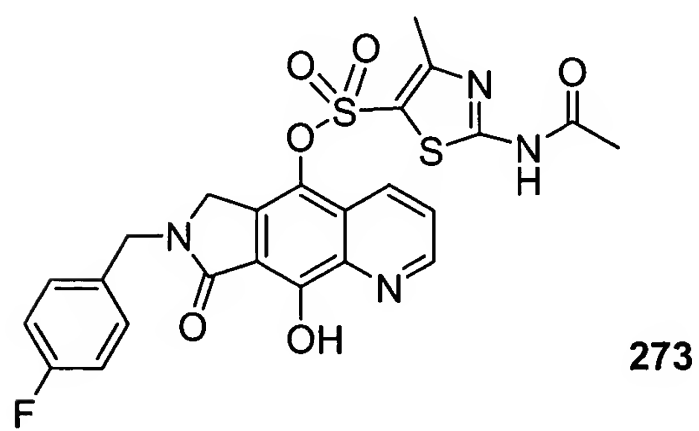
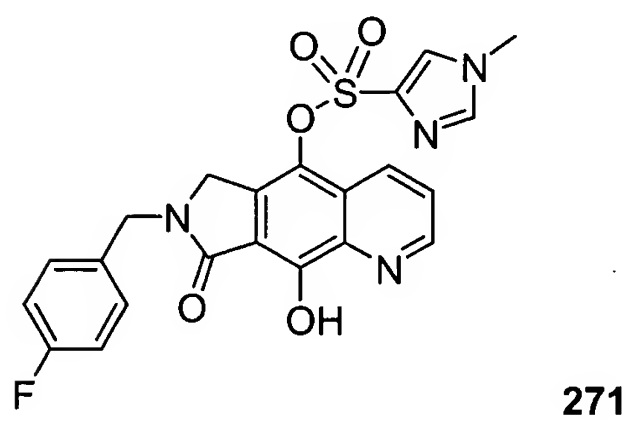
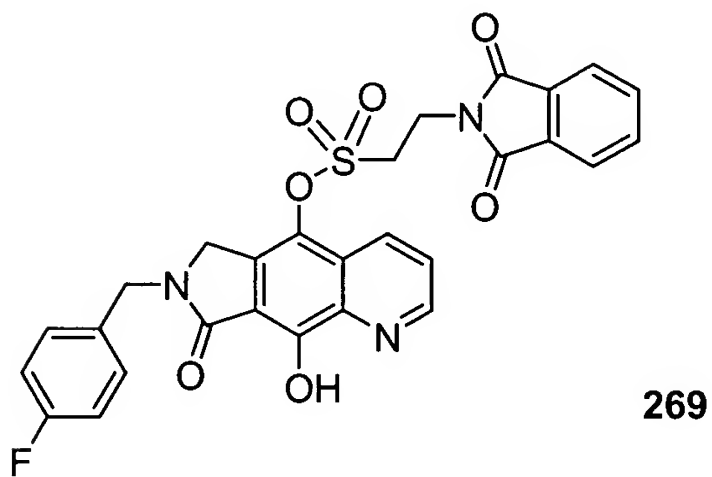
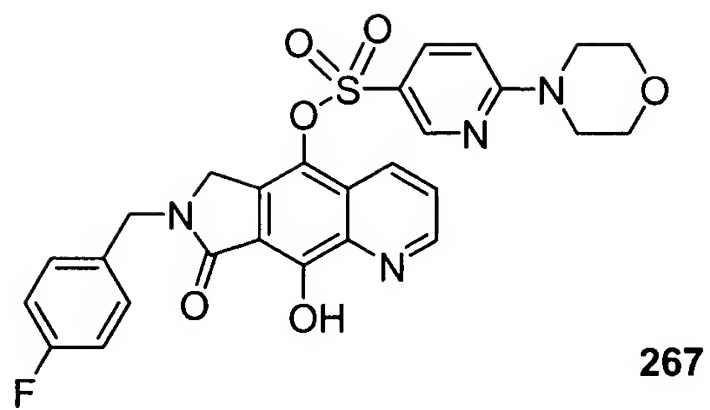
240

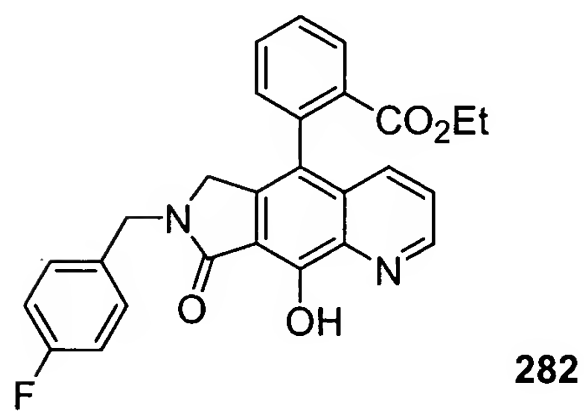
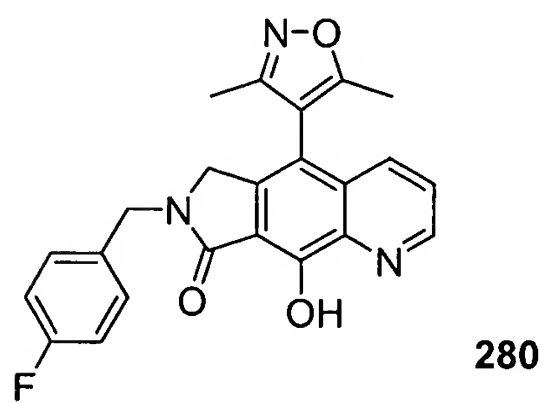
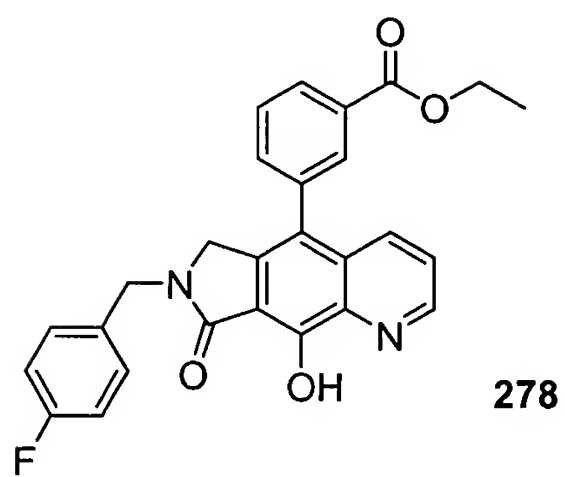
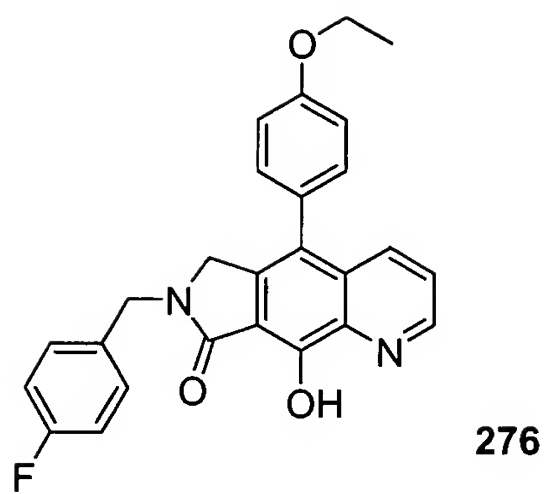


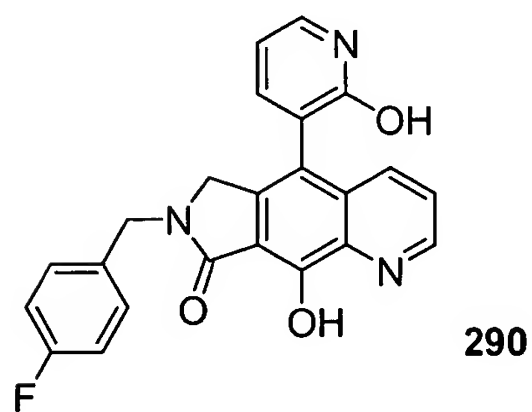
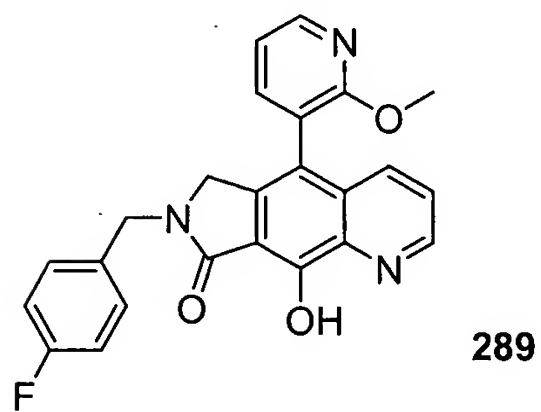
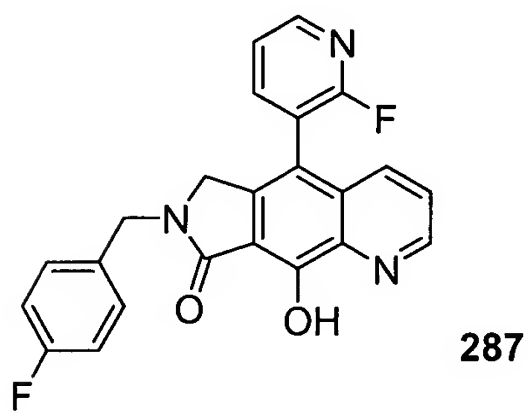
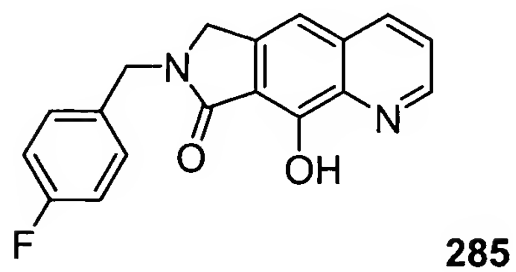
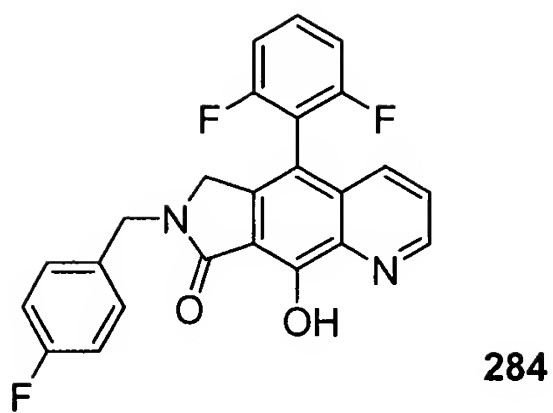
263

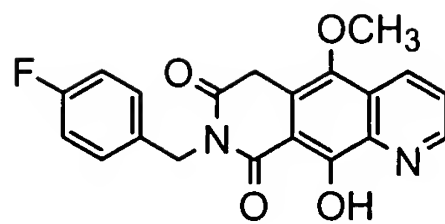


265

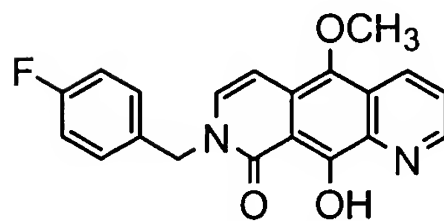




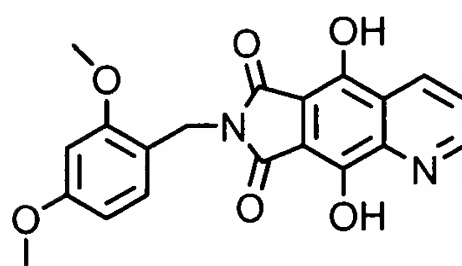




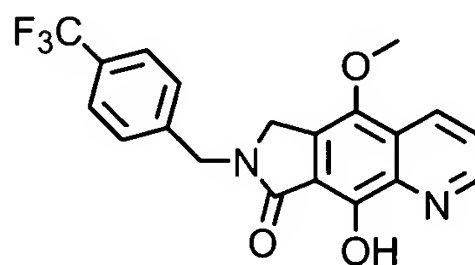
296



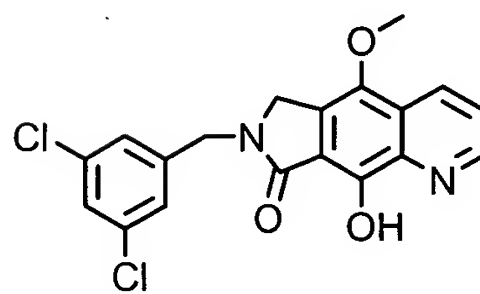
298



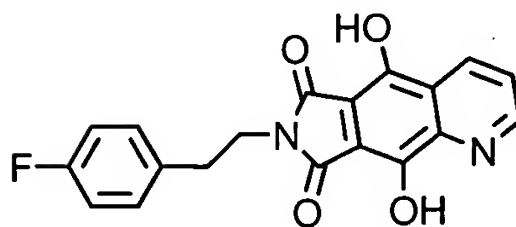
300



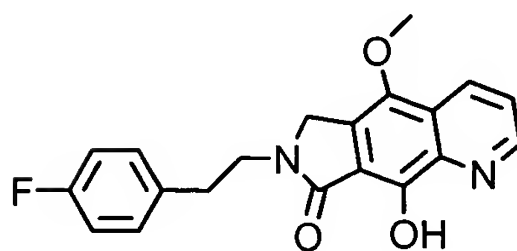
306



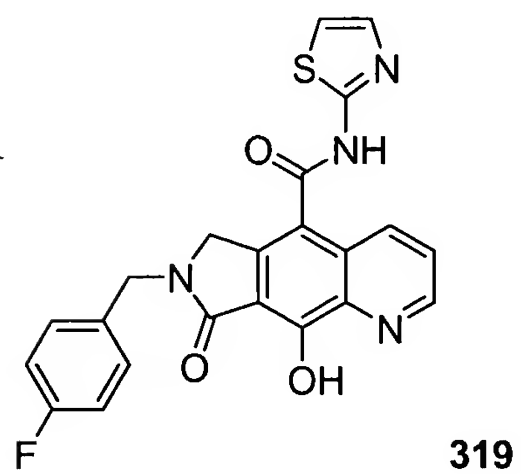
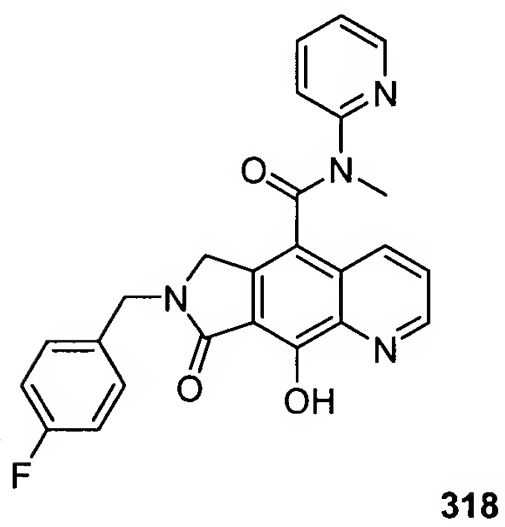
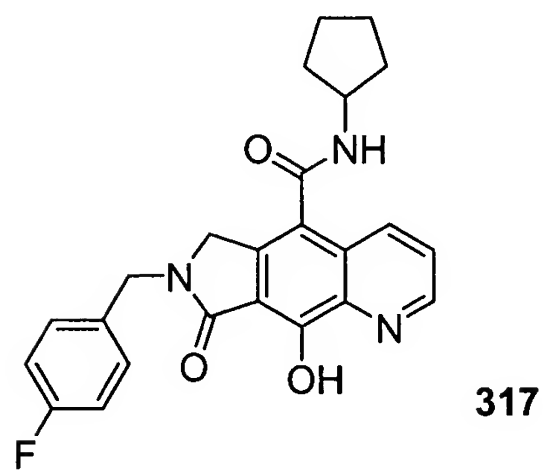
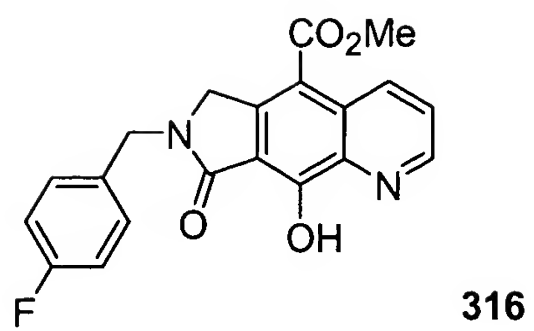
308

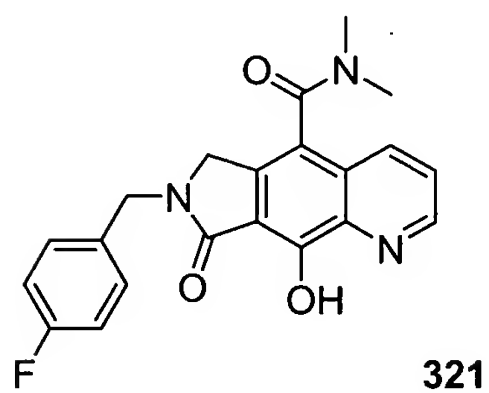
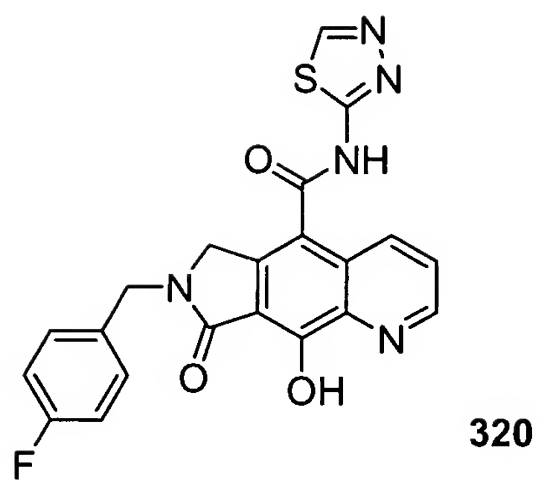


310

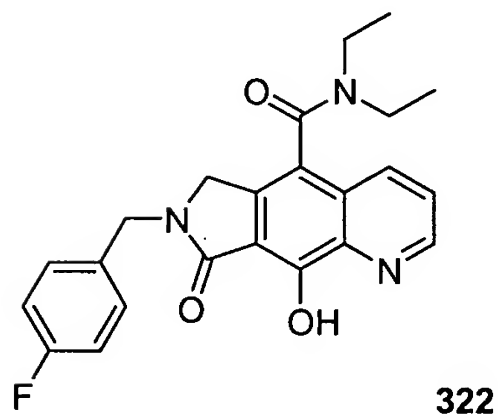


313

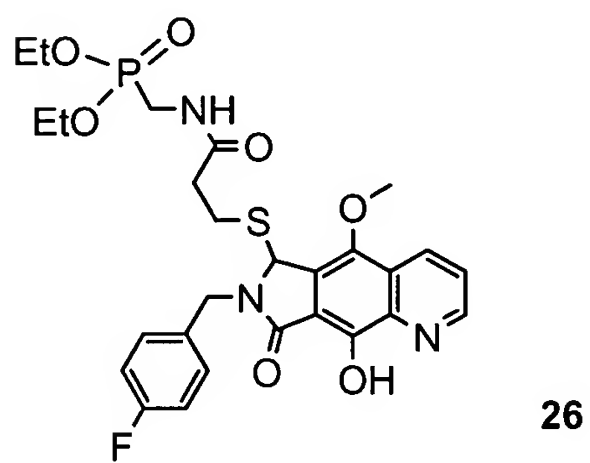


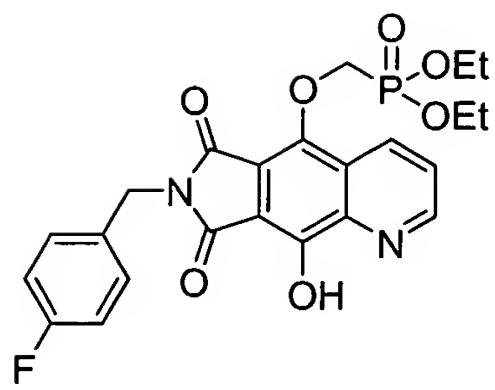


and

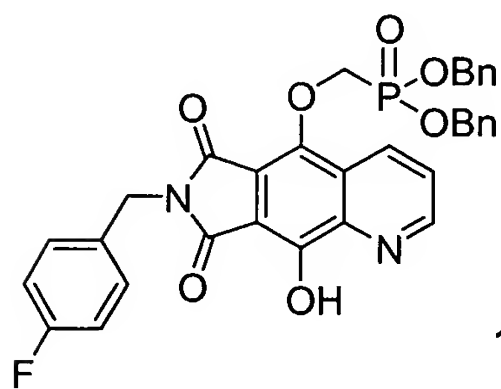


48. (Previously presented): A compound selected from the structures:

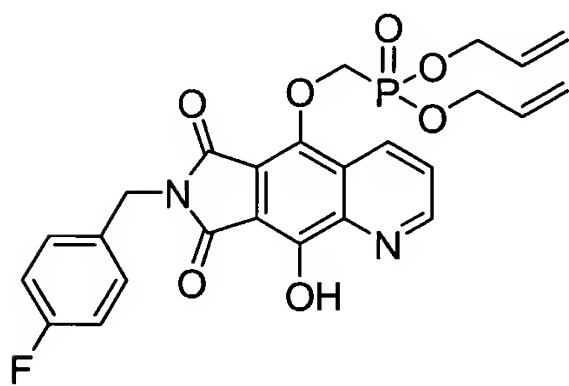




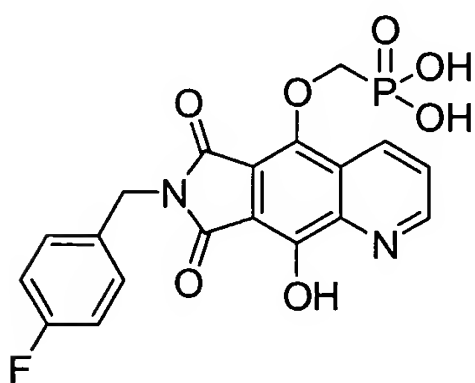
149



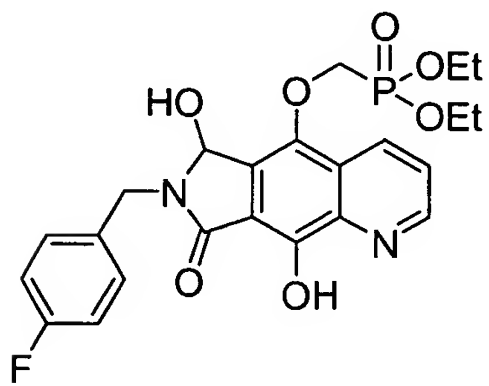
151



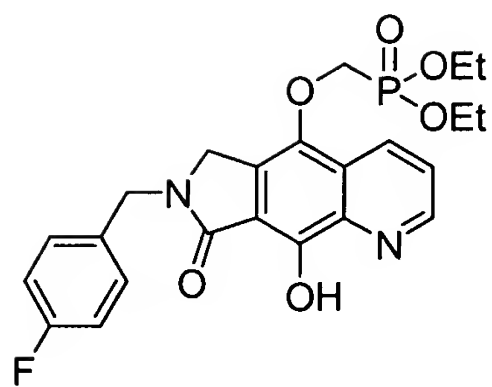
153



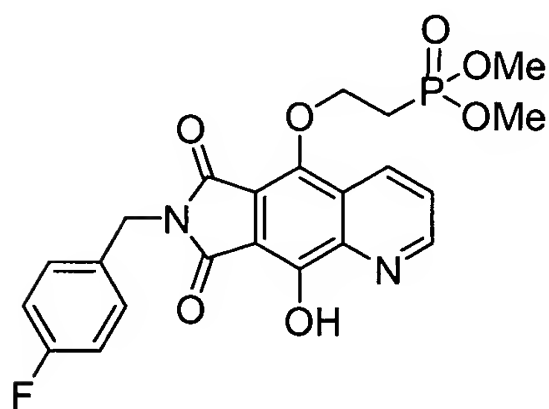
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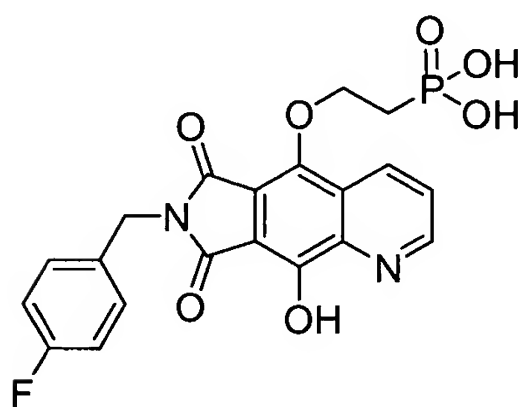
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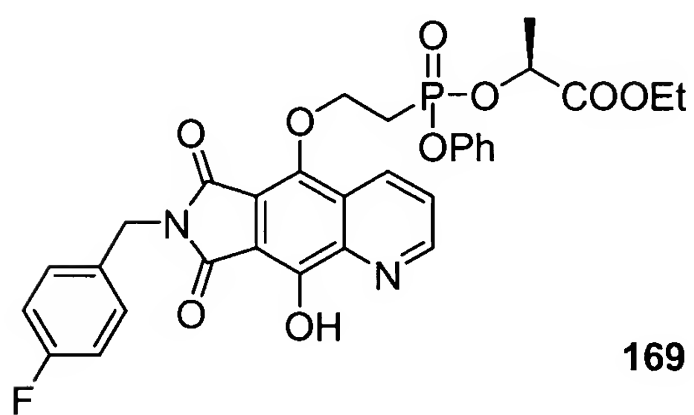
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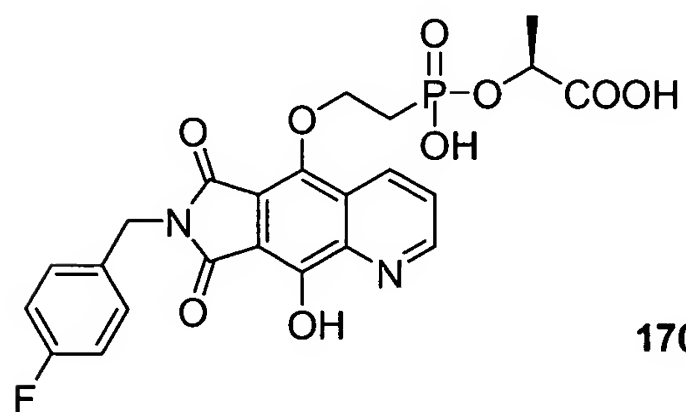
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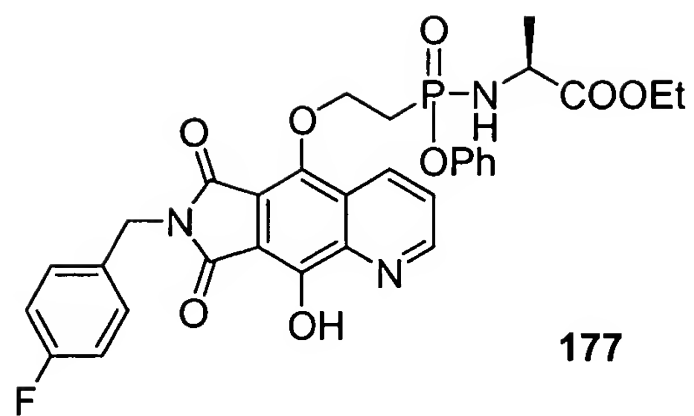
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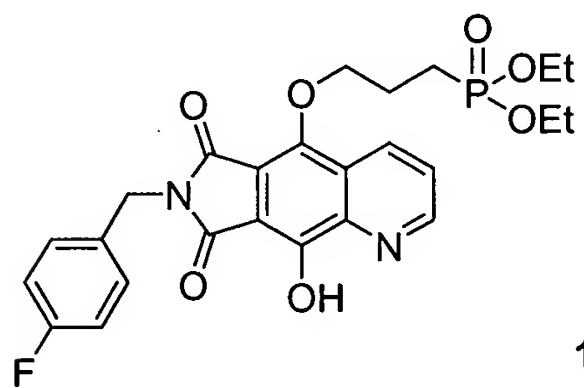
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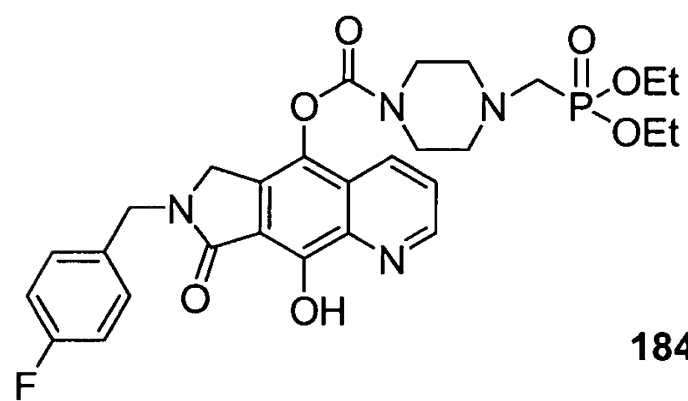
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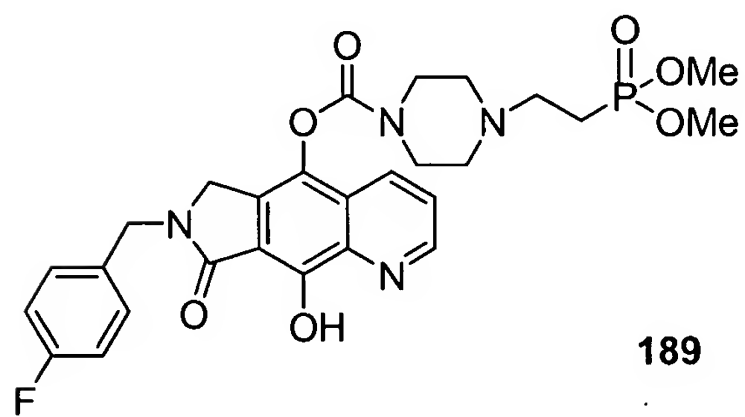
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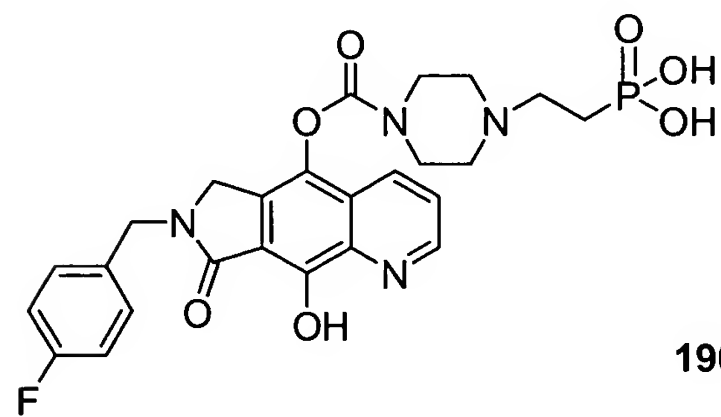
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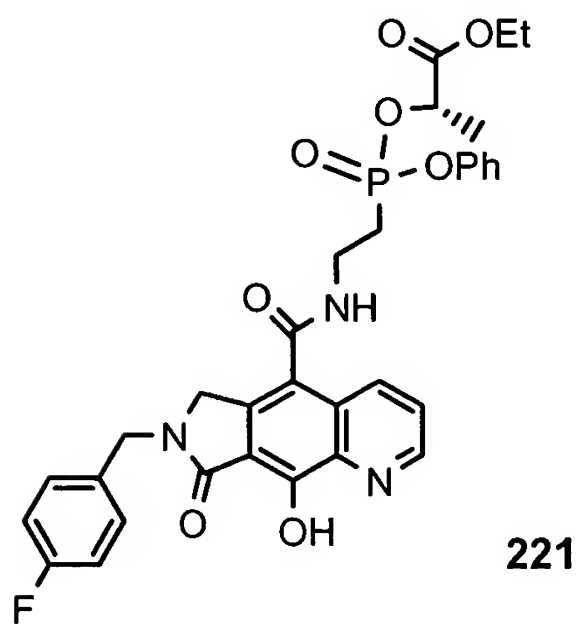
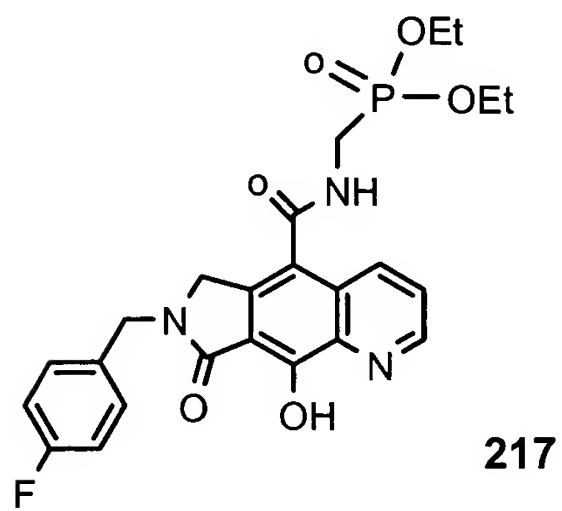
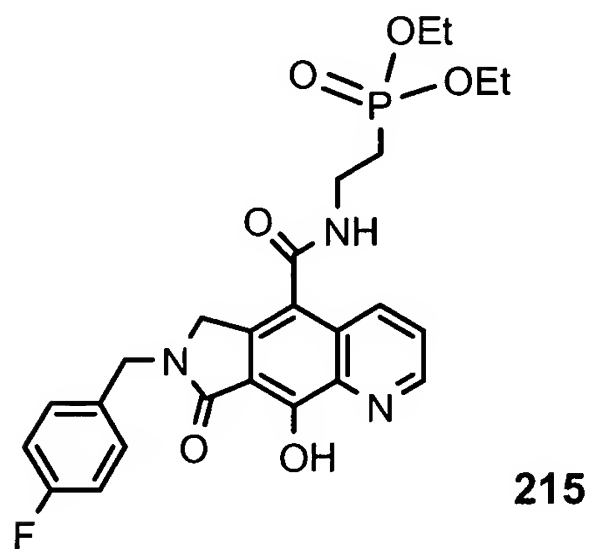
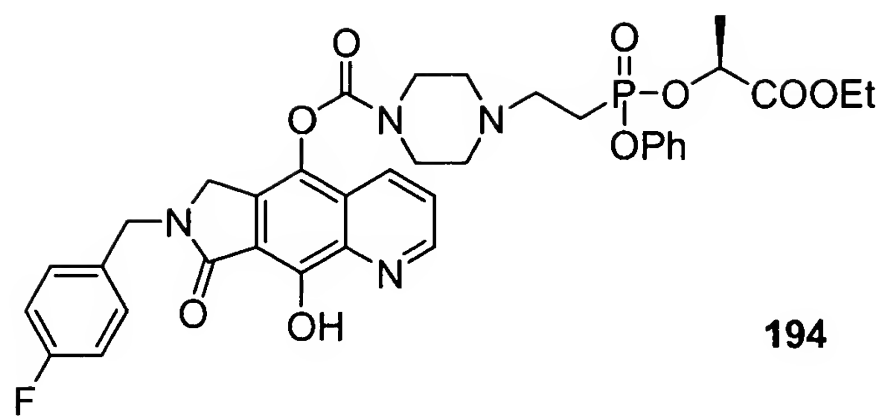
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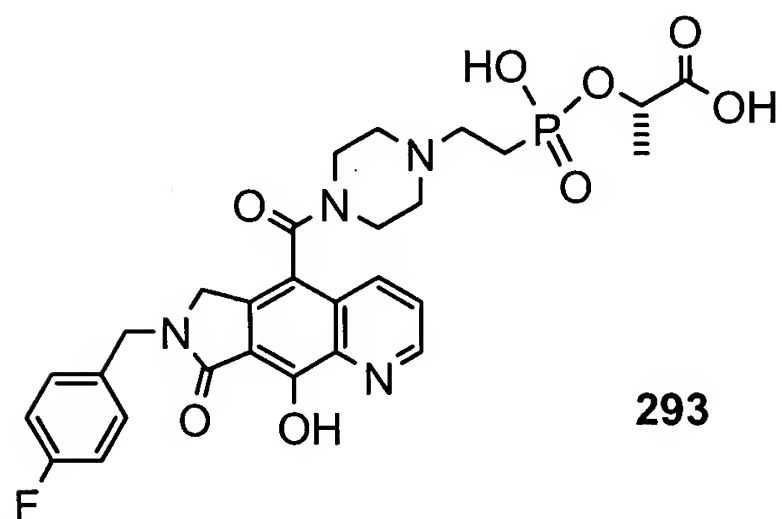
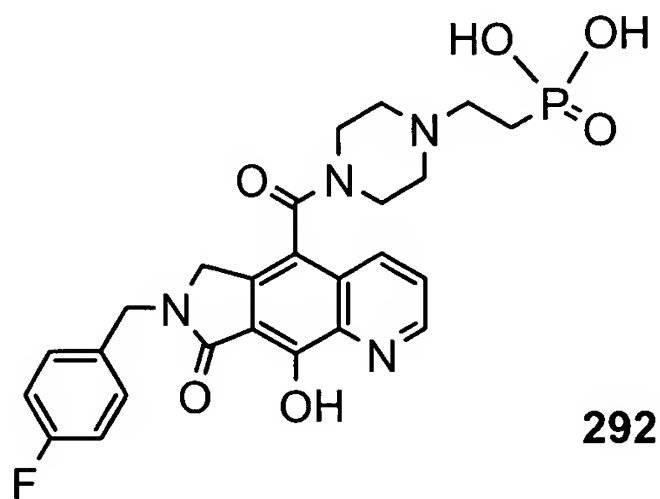
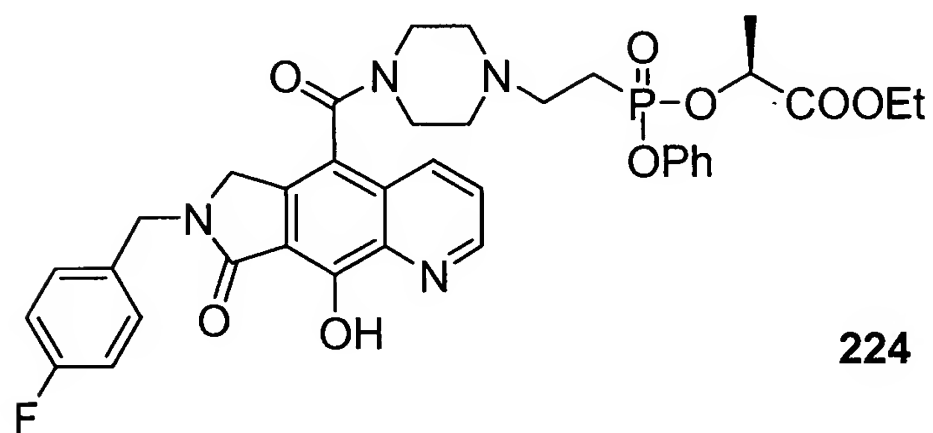
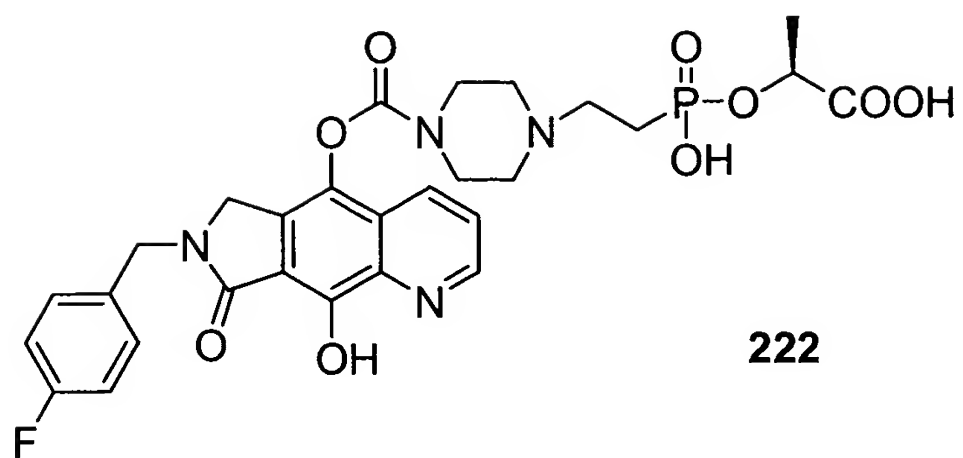


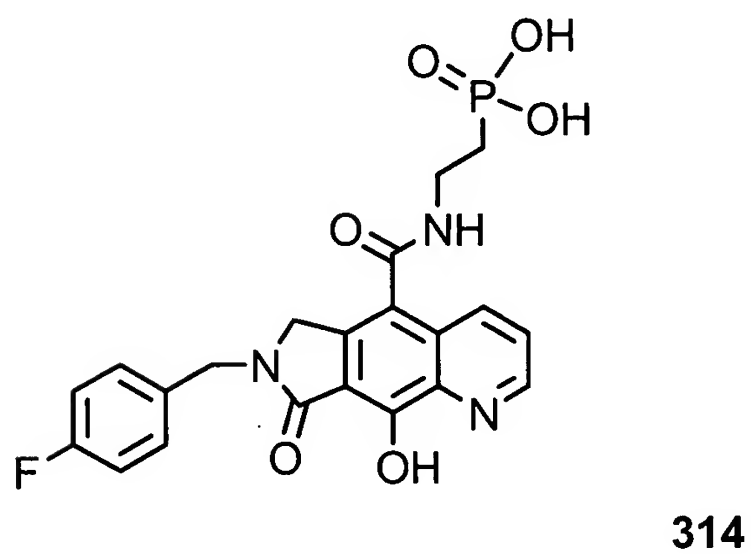
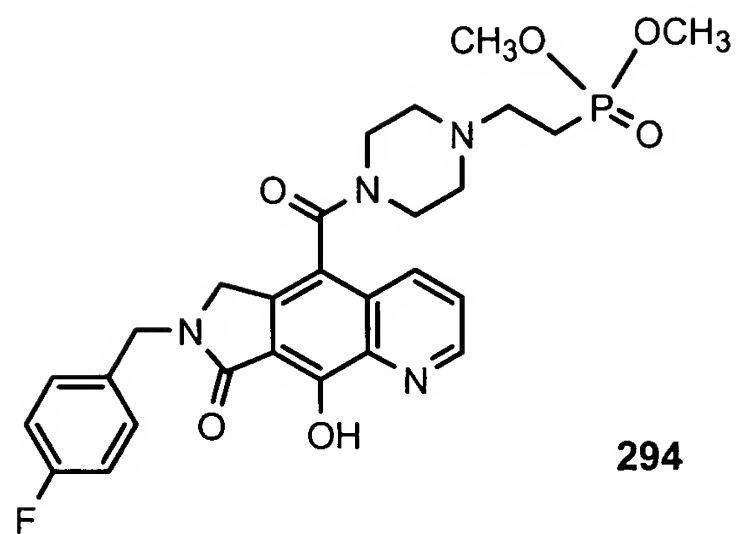
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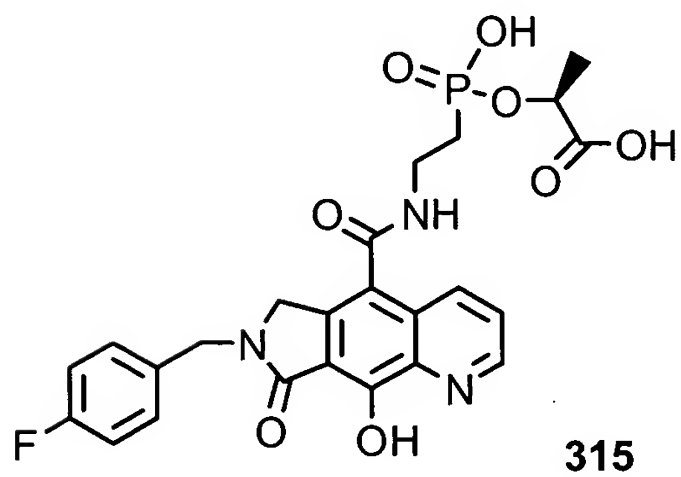
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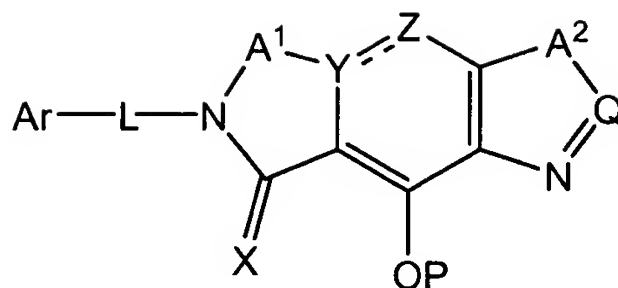




and



49. (Previously presented): A compound having the structure:

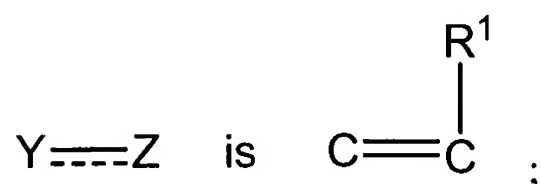


wherein:

A¹ is independently selected from C(R²)₂, CR²OR, CR²OC(=O)R, C(=O), C(=S), CR²SR, and C(=NR),

A² is independently selected from C(R²)₂-C(R³)₂, C(R²)=C(R³), and C(=O)C(R³)₂;

Q is CR⁴;



L is selected from a bond, O, S, S-S, S(=O), S(=O)₂, S(=O)₂NR, NR, N-OR, C₁-C₁₂ alkylene, C₁-C₁₂ substituted alkylene, C₂-C₁₂ alkenylene, C₂-C₁₂ substituted alkenylene, C₂-C₁₂ alkynylene, C₂-C₁₂ substituted alkynylene, C(=O)NH, OC(=O)NH, NHC(=O)NH, C(=O), C(=O)NH(CH₂)_n, or (CH₂CH₂O)_n, where n is optionally 1, 2, 3, 4, 5, or 6;

X is selected from O, S, NH, NR, N-OR, N-NR₂, N-CR₂OR and N-CR₂NR₂;

Ar is selected from (a) a C₃-C₁₂ carbocycle, C₃-C₁₂ substituted carbocycle, C₆-C₂₀ aryl, C₆-C₂₀ substituted aryl, C₂-C₂₀ heteroaryl, and C₂-C₂₀ substituted heteroaryl;

or (b) a saturated, unsaturated or aromatic ring or ring system having a mono- or bicyclic carbocycle or heterocycle containing 3 to 12 ring atoms;

R², R³ and R⁴ are each independently selected from H, F, Cl, Br, I, OH, -NH₂, -NH₃⁺, -NHR, -NR₂, -NR₃⁺, C₁-C₈ alkylhalide, carboxylate, sulfate, sulfamate, sulfonate, 5-7 membered ring sultam, C₁-C₈ alkylsulfonate, C₁-C₈ alkylamino, 4-dialkylaminopyridinium, C₁-C₈ alkylhydroxyl, C₁-C₈ alkylthiol, -SO₂R, -SO₂Ar, -SOAr, -SAr, -SO₂NR₂, -SOR, -CO₂R, -C(=O)NR₂, 5-7 membered ring lactam, 5-7 membered ring lactone, -CN, -N₃, -NO₂, C₁-C₈ alkoxy, C₁-C₈ trifluoroalkyl, C₁-C₈ alkyl, C₁-C₈ substituted alkyl, C₃-C₁₂ carbocycle, C₃-C₁₂ substituted carbocycle, C₆-C₂₀ aryl, C₆-C₂₀ substituted aryl, C₂-C₂₀ heteroaryl, and C₂-C₂₀ substituted heteroaryl, polyethyleneoxy, phosphonate, and phosphate;

when taken together on a single carbon, two R^2 or two R^3 may form a spiro ring;

R^1 is independently selected from CR_3 , $NRSO_2R$, $OC(=O)NR_2$, $OC(=O)R$, SR , H , F , Cl , Br , I , OH , $-NH_2$, $-NH_3^+$, $-NHR$, $-NR_2$, $-NR_3^+$, C_1-C_8 alkylhalide, carboxylate, sulfate, sulfamate, sulfonate, 5-7 membered ring sultam, C_1-C_8 alkylsulfonate, C_1-C_8 alkylamino, 4-dialkylaminopyridinium, C_1-C_8 alkylhydroxyl, C_1-C_8 alkylthiol, $-SO_2R$, $-SO_2Ar$, $-SOAr$, $-SAr$, $-SO_2NR_2$, $-SOR$, $-CO_2R$, $-C(=O)NR_2$, 5-7 membered ring lactam, 5-7 membered ring lactone, $-CN$, $-N_3$, $-NO_2$, C_1-C_8 alkoxy, C_1-C_8 trifluoroalkyl, C_1-C_8 alkyl, C_1-C_8 substituted alkyl, C_3-C_{12} carbocycle, C_3-C_{12} substituted carbocycle, C_6-C_{20} aryl, C_6-C_{20} substituted aryl, C_2-C_{20} heteroaryl, and C_2-C_{20} substituted heteroaryl, polyethyleneoxy, phosphonate, and phosphate;

R^1 is independently selected from CR_3 , $NRSO_2R$, $OC(=O)OR$, $OC(=O)NR_2$, $OC(+O)R$, SR , H , F , Cl , Br , I , OH , $-NH_2$, $-NH_3^+$, $-NHR$, $-NR_2$, $-NR_3^+$, C_1-C_8 alkylhalide, carboxylate, sulfate, sulfamate, sulfonate, 5-7 membered ring sultam, C_1-C_8 alkylsulfonate, C_1-C_8 alkylamino, 4-dialkylaminopyridinium, C_1-C_8 alkylhydroxyl, C_1-C_8 alkylthiol, $-SO_2R$, $-SO_2Ar$, $-SOAr$, $-SAr$, $-SO_2NR_2$, $-SOR$, $-CO_2R$, $-C(=O)NR_2$, 5-7 membered ring lactam, 5-7 membered ring lactone, $-CN$, $-N_3$, $-NO_2$, C_1-C_8 alkoxy, C_1-C_8 trifluoroalkyl, C_1-C_8 alkyl, C_1-C_8 substituted alkyl, C_3-C_{12} carbocycle, C_3-C_{12} substituted carbocycle, C_6-C_{20} aryl, C_6-C_{20} substituted aryl, C_2-C_{20} heteroaryl, and C_2-C_{20} substituted heteroaryl, polyethyleneoxy, phosphonate, and phosphate;

R is independently selected from H , C_1-C_8 alkyl, C_1-C_8 substituted alkyl, C_6-C_{20} aryl, C_6-C_{20} substituted aryl, C_2-C_{20} heteroaryl, and C_2-C_{20} substituted heteroaryl, polyethyleneoxy, phosphonate, and phosphate;

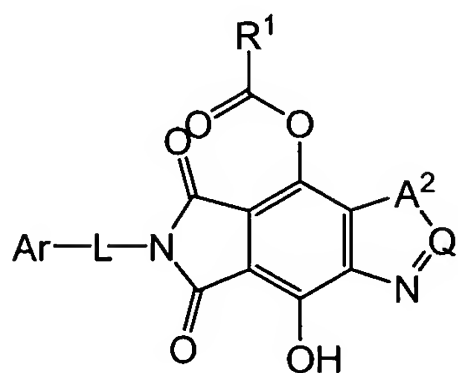
R^{X2} is independently selected from H , C_1-C_8 alkyl, C_1-C_8 substituted alkyl, C_6-C_{20} aryl, C_6-C_{20} substituted aryl, C_2-C_{20} heteroaryl, and C_2-C_{20} substituted heteroaryl, polyethyleneoxy, phosphonate, and phosphate;

and the tautomers, salts, solvates, resolved enantiomers and purified diastereomers thereof;

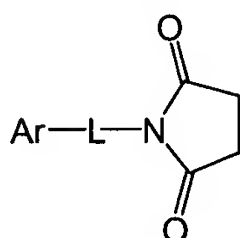
with the proviso that when $Y=Z$ is $C=C(OH)$, X is O , A^1 is $C(=O)$, A^2 is $C(R^2)=C(R^3)$, and Q is CH , then L is not a bond; and

P is a protecting group selected from benzyhydryl ($CHPh_2$), trialkylsilyl (R_3Si), 2-trimethylsiloxyethyl, alkoxymethyl (CH_2OR), and ester ($C(=O)R$).

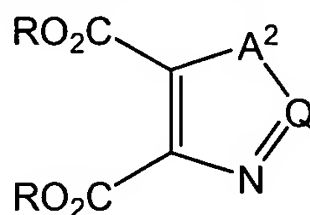
50. (Previously presented): A process for preparation of a compound having the structure:



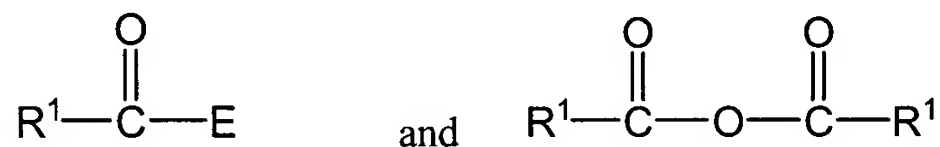
comprising reacting a succinimide compound having the structure:



with a heterocyclic compound having the structure:



and reacting with an acylation reagent having a formula selected from:



wherein:

A² is selected from C(R²)₂-C(R³)₂, C(R²)=C(R³), and C(=O)C(R³)₂;

Q is CR⁴;

L is selected from a bond, O, S, NR, N-OR, C₁-C₁₂ alkyldiyl, C₁-C₁₂ substituted alkyldiyl, C(=O)NH, C(=O), S(=O), S(=O)₂, C(=O)NH(CH₂)_n, and (CH₂CH₂O)_n, where n ranges from 1 to 6;

Ar is selected from (a) a C₆-C₂₀ aryl, C₆-C₂₀ substituted aryl, C₂-C₂₀ heteroaryl, and C₂-C₂₀ substituted heteroaryl or (b) a saturated, unsaturated or aromatic ring or ring system having a mono- or bicyclic carbocycle or heterocycle containing 3 to 12 ring atoms;

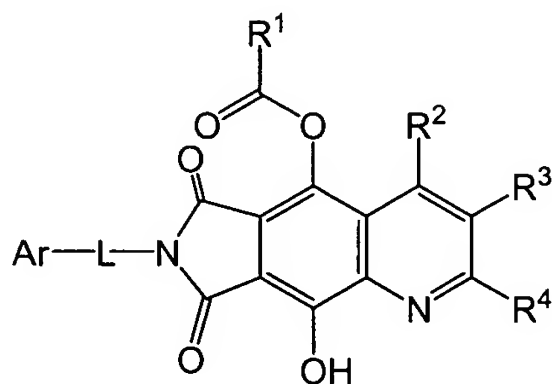
R¹ is selected from R, OR, NR₂, NHR, NHSO₂R, and NRSO₂R;

E is selected from Cl, imidazole, and hydroxybenzotriazole;

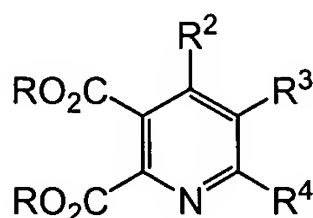
R^2 , R^3 and R^4 are each independently selected from H, F, Cl, Br, I, OH, $-NH_2$, $-NH_3^+$, $-NHR$, $-NR_2$, $-NR_3^+$, C_1-C_8 alkylhalide, carboxylate, sulfate, sulfamate, sulfonate, 5-7 membered ring sultam, C_1-C_8 alkylsulfonate, C_1-C_8 alkylamino, 4-dialkylaminopyridinium, C_1-C_8 alkylhydroxyl, C_1-C_8 alkylthiol, $-SO_2R$, $-SO_2Ar$, $-SOAr$, $-SAr$, $-SO_2NR_2$, $-SOR$, $-CO_2R$, $-C(=O)NR_2$, 5-7 membered ring lactam, 5-7 membered ring lactone, $-CN$, $-N_3$, $-NO_2$, C_1-C_8 alkoxy, C_1-C_8 trifluoroalkyl, C_1-C_8 alkyl, C_1-C_8 substituted alkyl, C_3-C_{12} carbocycle, C_3-C_{12} substituted carbocycle, C_6-C_{20} aryl, C_6-C_{20} substituted aryl, C_2-C_{20} heteroaryl, and C_2-C_{20} substituted heteroaryl, polyethyleneoxy, phosphonate, phosphate, and a prodrug moiety; and

R is selected from C_1-C_6 alkyl, C_1-C_6 substituted alkyl, C_6-C_{20} aryl, C_6-C_{20} substituted aryl, C_2-C_{20} heteroaryl, C_2-C_{20} substituted heteroaryl, polyethyleneoxy, phosphonate, phosphate, and a prodrug moiety.

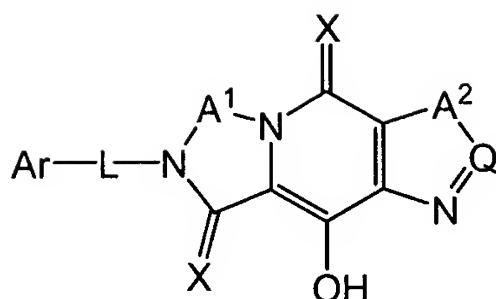
51. (Previously presented): The process of claim 50 for preparation of a compound having the structure:



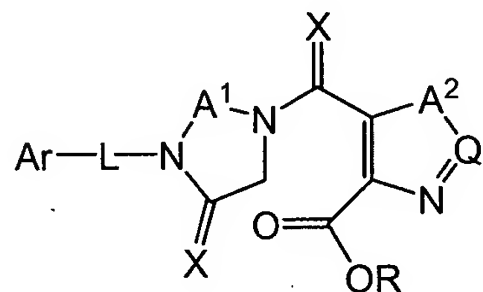
wherein the heterocyclic compound has the structure:



52. (Previously presented): A process for preparation of a compound having the structure:



comprising reacting a compound having the structure:



with a basic reagent comprising hydroxide, an alkoxide or an amine;

wherein:

A^1 is independently selected from $C(R^2)_2$, CR^2OR , $CR^2OC(=O)R$, $C(=O)$, $C(=S)$, CR^2SR , and $C(=NR)$,

A^2 is independently selected from $C(R^2)_2-C(R^3)_2$, $C(R^2)=C(R^3)$, and $C(=O)C(R^3)_2$;

Q is CR^4 ;

X is selected from O , S , NH , NR , $N-OR$, $N-NR_2$, $N-CR_2OR$ and $N-CR_2NR_2$;

L is selected from a bond, O , S , NR , $N-OR$, C_1-C_{12} alkyldiyl, C_1-C_{12} substituted alkyldiyl, $C(=O)NH$, $C(=O)$, $S(=O)$, $S(=O)_2$, $C(=O)NH(CH_2)_n$, and $(CH_2CH_2O)_n$, where n ranges from 1 to 6;

Ar is selected from (a) a C_6-C_{20} aryl, C_6-C_{20} substituted aryl, C_2-C_{20} heteroaryl, and C_2-C_{20} substituted heteroaryl;

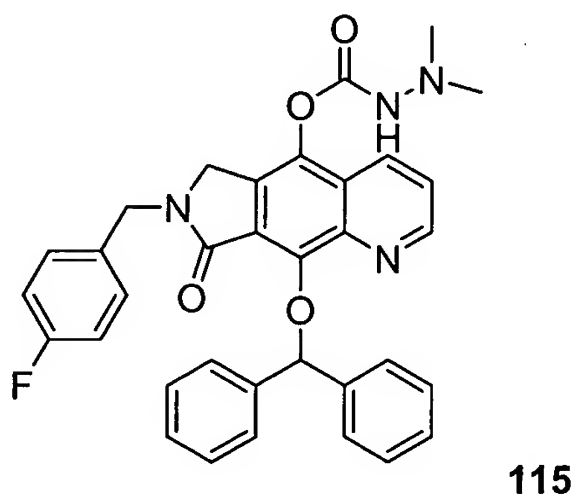
or (b) a saturated, unsaturated or aromatic ring or ring system having a mono- or bicyclic carbocycle or heterocycle containing 3 to 12 ring atoms;

R^2 , R^3 and R^4 are each independently selected from H , F , Cl , Br , I , OH , $-NH_2$, $-NH_3^+$, $-NHR$, $-NR_2$, $-NR_3^+$, C_1-C_8 alkylhalide, carboxylate, sulfate, sulfamate, sulfonate, 5-7 membered ring sultam, C_1-C_8 alkylsulfonate, C_1-C_8 alkylamino, 4-dialkylaminopyridinium, C_1-C_8 alkylhydroxyl, C_1-C_8 alkylthiol, $-SO_2R$, $-SO_2Ar$, $-SOAr$, $-SAr$, $-SO_2NR_2$, $-SOR$, $-CO_2R$, $-C(=O)NR_2$, 5-7 membered ring lactam, 5-7 membered ring lactone, $-CN$, $-N_3$, $-NO_2$, C_1-C_8 alkoxy, C_1-C_8 trifluoroalkyl, C_1-C_8 alkyl, C_1-C_8 substituted alkyl, C_3-C_{12} carbocycle, C_3-C_{12}

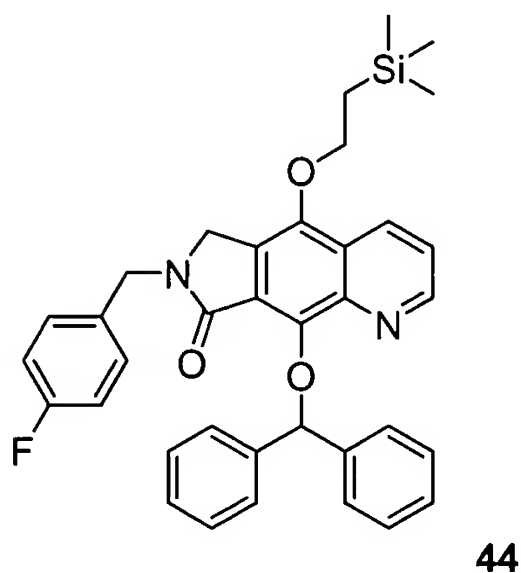
substituted carbocycle, C₆-C₂₀ aryl, C₆-C₂₀ substituted aryl, C₂-C₂₀ heteroaryl, and C₂-C₂₀ substituted heteroaryl, polyethyleneoxy, phosphonate, phosphate, and a prodrug moiety; and

R is selected from C₁-C₆ alkyl, C₁-C₆ substituted alkyl, C₆-C₂₀ aryl, C₆-C₂₀ substituted aryl, C₂-C₂₀ heteroaryl, C₂-C₂₀ substituted heteroaryl, polyethyleneoxy, phosphonate, phosphate, and a prodrug moiety.

53. (Previously presented): A process for preparation of a compound having structure 115:



comprising reacting a compound having the structure 44:



with tetrabutylammonium fluoride to form a desilylated intermediate; and
reacting the desilylated intermediate with triphosgene (bis(trichloromethyl) carbonate),
followed by dimethylhydrazine to form structure 115.

54. (Previously presented): A compound of claim 1 substituted with phosphonate and capable of accumulating in human PBMC.

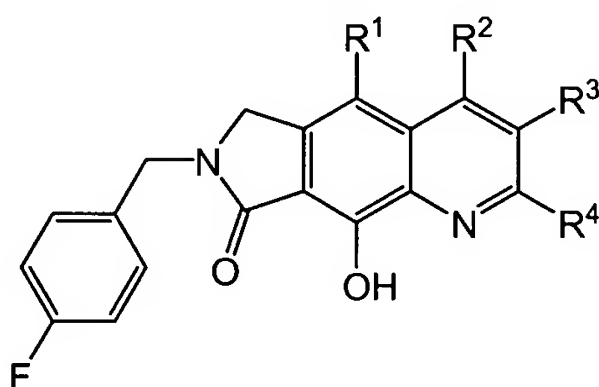
55. (Previously presented): The compound of claim 54 wherein the intracellular half-life of the compound or an intracellular metabolite of the compound in human PBMC is increased by at least about 50% when compared to the analog of the compound not having the phosphonate.
56. (canceled)
57. (canceled)
58. (Previously presented): The compound of claim 55 wherein the half-life is improved by at least about 100%.
59. (canceled)
60. (canceled)
61. (canceled)
62. (canceled)
63. (Previously presented): A pharmaceutical composition comprising a therapeutically effective amount of a compound of claim 1 and a pharmaceutically acceptable carrier.
64. (Withdrawn): The pharmaceutical composition of claim 63 further comprising a therapeutically effective amount of an AIDS treatment agent selected from an HIV inhibitor agent, an anti-infective agent, and an immunomodulator.
65. (Withdrawn): The pharmaceutical composition of claim 64 wherein the HIV inhibitor agent is an HIV-protease inhibitor.
66. (Withdrawn): The composition of claim 64 wherein the HIV inhibitor agent is a nucleoside reverse transcriptase inhibitor.
67. (Withdrawn): The composition of claim 64 wherein the HIV inhibitor agent is a non-nucleoside reverse transcriptase inhibitor.
68. (Withdrawn): A process for making a pharmaceutical composition comprising combining a compound of claim 1 and a pharmaceutically acceptable carrier.

69. (Withdrawn): A method of inhibiting HIV integrase, comprising the administration to a mammal in need of such treatment of a therapeutically effective amount of a compound of claim 1.

70. (Withdrawn): A method of treating infection by HIV, or of treating AIDS or ARC, comprising administration to a mammal in need of such treatment of a therapeutically effective amount of a compound of claim 1.

71 – 79 (canceled)

80. (Previously presented): A compound having the structure:



wherein R¹ is selected from R, OR, NR₂, NHR, NHSO₂R and NRSO₂R;

R², R³ and R⁴ are each independently selected from H, F, Cl, Br, I, OH, -NH₂, -NH₃⁺, -NHR, -NR₂, -NR₃⁺, C₁-C₈ alkylhalide, carboxylate, sulfate, sulfamate, sulfonate, 5-7 membered ring sultam, C₁-C₈ alkylsulfonate, C₁-C₈ alkylamino, 4-dialkylaminopyridinium, C₁-C₈ alkylhydroxyl, C₁-C₈ alkylthiol, -SO₂R, -SO₂Ar, -SOAr, -SAr, -SO₂NR₂, -SOR, -CO₂R, -C(=O)NR₂, 5-7 membered ring lactam, 5-7 membered ring lactone, -CN, -N₃, -NO₂, C₁-C₈ alkoxy, C₁-C₈ trifluoroalkyl, C₁-C₈ alkyl, C₁-C₈ substituted alkyl, C₃-C₁₂ carbocycle, C₃-C₁₂ substituted carbocycle, C₆-C₂₀ aryl, C₆-C₂₀ substituted aryl, C₂-C₂₀ heteroaryl, and C₂-C₂₀ substituted heteroaryl, polyethyleneoxy, phosphonate, phosphate, and a prodrug moiety; and

R is independently selected from H, C₁-C₈ alkyl, C₁-C₈ substituted alkyl, C₆-C₂₀ aryl, C₆-C₂₀ substituted aryl, C₂-C₂₀ heteroaryl, and C₂-C₂₀ substituted heteroaryl, polyethyleneoxy, phosphonate, phosphate, and a prodrug moiety;

and the tautomers, salts, solvates, resolved enantiomers and purified diastereomers thereof.

81. (Withdrawn): Method of treating a disorder affecting white blood cells, comprising:
administering a compound of claim 1 comprising phosphonate prodrug to a patient in need of white-blood-cell targeting.
82. (Withdrawn): Method of targeting a compound to white blood cells, comprising:
selecting a compound of claim 1 having a desired pharmaceutical activity and having a first structure;
modifying said first structure by replacing one or more atom of said first structure with an organic substituent comprising a phosphonate group or incipient phosphonate group to provide a compound having a second structure.
83. (Withdrawn): A method of manufacturing an HIV inhibitor compound having both selectivity for white blood cells and a desired pharmaceutical activity, comprising:
chemically synthesizing a first molecule of claim 1 having a first structure containing a phosphonate or precursor phosphonate group, wherein said first structure differs from a second structure of a compound known to have said desired pharmaceutical activity by having at least one hydrogen atom of said second structure replaced by an organic substituent comprising a phosphonate group or incipient phosphonate group.
84. (Withdrawn): The method of claim 83, wherein said first molecule is synthesized by a series of chemical reactions in which a hydrogen of said second structure is replaced by said organic substituent.
85. (Withdrawn): The method of claim 83, wherein said first molecule is synthesized by a series of chemical reactions that never includes a molecule of said second structure.
86. (Withdrawn): Method of accumulating an HIV integrase inhibitor compound inside a white blood cell, comprising:

administering to a sample a composition comprising a compound of claim 1.

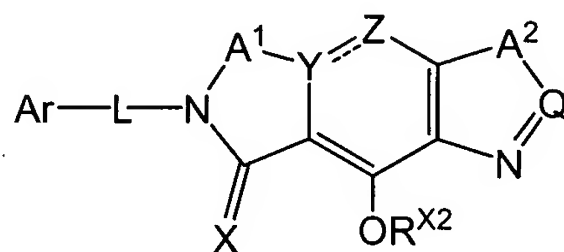
87. (Withdrawn): The method of claim 85 wherein said sample is a patient.

88. (Withdrawn): The method of claim 83, wherein said compound has a chemical structure A-B, wherein (a) a compound having structure A-H has HIV integrase inhibitor activity and (b) substructure B comprises a phosphonate group or a precursor phosphonate group.

89. (Withdrawn): Method of increasing half-life of an HIV integrase inhibitor compound, comprising:

replacing at least one hydrogen atom or organic radical of a compound of claim 1 by an organic substituent comprising a phosphonate group or incipient phosphonate.

90. (New): A compound having the structure:

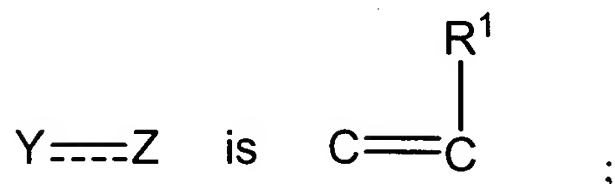


wherein:

A¹ is independently selected from C(R²)₂, CR²OR, CR²OC(=O)R, C(=O), C(=S), CR²SR, and C(=NR),

A² is independently selected from C(R²)₂-C(R³)₂, C(R²)=C(R³) and C(=O)C(R³)₂;

Q is CR⁴;



L is selected from a bond, O, S, S-S, S(=O), S(=O)₂, S(=O)₂NR, NR, N-OR, C₁-C₁₂ alkylene, C₁-C₁₂ substituted alkylene, C₂-C₁₂ alkenylene, C₂-C₁₂ substituted alkenylene, C₂-C₁₂

alkynylene, C₂–C₁₂ substituted alkynylene, C(=O)NH, OC(=O)NH, NHC(=O)NH, C(=O), C(=O)NH(CH₂)_n, or (CH₂CH₂O)_n, where n is optionally 1, 2, 3, 4, 5, or 6;

X is selected from O, S, NH, NR, N–OR, N–NR₂, N–CR₂OR and N–CR₂NR₂;

Ar is selected from (a) a C₃–C₁₂ carbocycle, C₃–C₁₂ substituted carbocycle, C₆–C₂₀ aryl, C₆–C₂₀ substituted aryl, C₂–C₂₀ heteroaryl, and C₂–C₂₀ substituted heteroaryl;

or (b) a saturated, unsaturated or aromatic ring or ring system having a mono- or bicyclic carbocycle or heterocycle containing 3 to 12 ring atoms;

R², R³ and R⁴ are each independently selected from H, F, Cl, Br, I, OH, –NH₂, –NH₃⁺, –NHR, –NR₂, –NR₃⁺, C₁–C₈ alkylhalide, carboxylate, sulfate, sulfamate, sulfonate, 5-7 membered ring sultam, C₁–C₈ alkylsulfonate, C₁–C₈ alkylamino, 4-dialkylaminopyridinium, C₁–C₈ alkylhydroxyl, C₁–C₈ alkylthiol, –SO₂R, –SO₂Ar, –SOAr, –SAr, –SO₂NR₂, –SOR, –CO₂R, –C(=O)NR₂, 5-7 membered ring lactam, 5-7 membered ring lactone, –CN, –N₃, –NO₂, C₁–C₈ alkoxy, C₁–C₈ trifluoroalkyl, C₁–C₈ alkyl, C₁–C₈ substituted alkyl, C₃–C₁₂ carbocycle, C₃–C₁₂ substituted carbocycle, C₆–C₂₀ aryl, C₆–C₂₀ substituted aryl, C₂–C₂₀ heteroaryl, and C₂–C₂₀ substituted heteroaryl, polyethyleneoxy, phosphonate, and phosphate;

when taken together on a single carbon, two R² or two R³ may form a spiro ring;

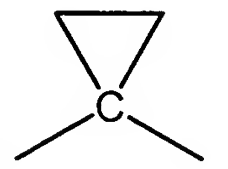
R¹ is independently selected from CR₃, NRSO₂R, OC(=O)NR₂, OC(=O)R, SR, H, F, Cl, Br, I, OH, –NH₂, –NH₃⁺, –NHR, –NR₂, –NR₃⁺, C₁–C₈ alkylhalide, carboxylate, sulfate, sulfamate, sulfonate, 5-7 membered ring sultam, C₁–C₈ alkylsulfonate, C₁–C₈ alkylamino, 4-dialkylaminopyridinium, C₁–C₈ alkylhydroxyl, C₁–C₈ alkylthiol, –SO₂R, –SO₂Ar, –SOAr, –SAr, –SO₂NR₂, –SOR, –CO₂R, –C(=O)NR₂, 5-7 membered ring lactam, 5-7 membered ring lactone, –CN, –N₃, –NO₂, C₁–C₈ alkoxy, C₁–C₈ trifluoroalkyl, C₁–C₈ alkyl, C₁–C₈ substituted alkyl, C₃–C₁₂ carbocycle, C₃–C₁₂ substituted carbocycle, C₆–C₂₀ aryl, C₆–C₂₀ substituted aryl, C₂–C₂₀ heteroaryl, and C₂–C₂₀ substituted heteroaryl, polyethyleneoxy, phosphonate, and phosphate;

R is independently selected from H, C₁–C₈ alkyl, C₁–C₈ substituted alkyl, C₆–C₂₀ aryl, C₆–C₂₀ substituted aryl, C₂–C₂₀ heteroaryl, and C₂–C₂₀ substituted heteroaryl, polyethyleneoxy, phosphonate, and phosphate;

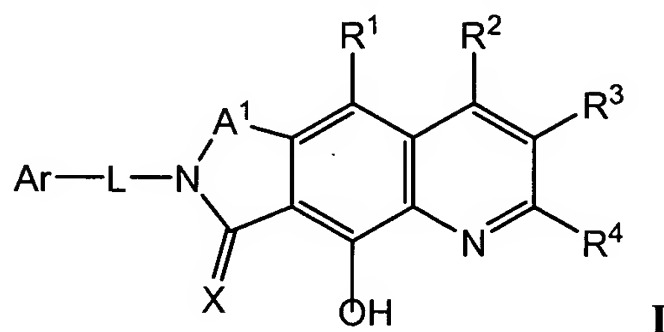
R^{X2} is independently selected from H, C₁–C₈ alkyl, C₁–C₈ substituted alkyl, C₆–C₂₀ aryl, C₆–C₂₀ substituted aryl, C₂–C₂₀ heteroaryl, and C₂–C₂₀ substituted heteroaryl, polyethyleneoxy, phosphonate, phosphate, and a protecting group;

and the tautomers, salts, solvates, resolved enantiomers and purified diastereomers thereof;
 with the proviso that when $Y=Z$ is $C=C(OH)$, X is O , A^1 is $C(=O)$, A^2 is $C(R^2)=C(R^3)$, and Q is CH , then L is not a bond.

91. (New): The compound of claim 90 wherein A^1 is CH_2 , or



92. (New): The compound of claim 90 having Formula I:

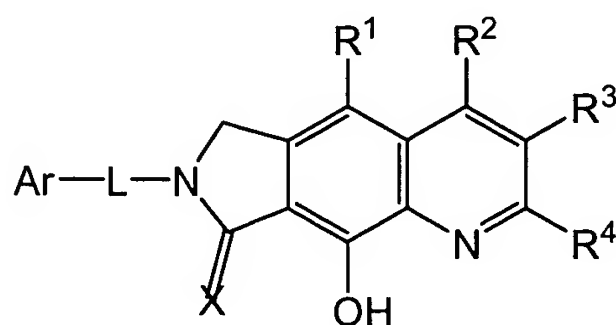


wherein X is O .

93. (New): The compound of claim 92 wherein L is CH_2 and Ar is substituted phenyl.

94. (New): The compound of claim 92 wherein A^1 is CH_2 and R^2 , R^3 and R^4 are each H .

95. (New): The compound of claim 90 having the Formula:



wherein X is O.

96. (New): The compound of claim 95 wherein L is CH₂ and Ar is substituted phenyl.
97. (New): The compound of claim 96 wherein R², R³, and R⁴ are each H.